SEQUENCE LISTING

<110> Norimasa; MATSUDA, Hikaru; SAWA, NAKAMURA, Yoshiki: TAKETANI. Satoshi; MIYAGAWA, Shigeru; YOSHIKAWA, Hideki; ANDO, Wataru <120> SCAFFOLD-FREE SELF-ORGANIZED 3D SYNTHETIC TISSUE <130> NKM001PCT <160> 30 <170> PatentIn version 3.2 <210> 1 <211> 6085 <212> DNA <213> Homo sapiens <220> <221> CDS <222> (115).. (5940) **<400>** 1 tacggctgcg agaagacgac agaagggggt cctgctttaa aaagctccaa gaactgtctc 60 actoccaggo tacatottot cacttgotaa caaggacoto tgagttcago agoc atg 117 Met 1 agt toa gac toa gaa ttg gct gtt ttt ggg gag gct gct cct ttc ctc 165 Ser Ser Asp Ser Glu Leu Ala Val Phe Gly Glu Ala Ala Pro Phe Leu 5 10 15 . cga aag tot gaa agg gag cgc att gag gcc cag aat agg ccc ttt gat 213

Arg Lys Ser Glu Arg Glu Arg Ile Glu Ala Gln Asn Arg Pro Phe Asp

 · gcc aaa aca tot gtc ttt gtg gcg gag ccc aaa gaa tcc ttt gtc aaa Ala Lys Thr Ser Val Phe Val Ala Glu Pro Lys Glu Ser Phe Val Lys ggg acc atc cag agc aga gaa gga gga aaa gtg acg gtg aag act gag Gly Thr Ile Gin Ser Arg Glu Gly Gly Lys Val Thr Val Lys Thr Glu gga gga gcg act ctg aca gtg aag gat gat cag gtc ttc ccc atg aac Gly Gly Ala Thr Leu Thr Val Lys Asp Asp Gln Val Phe Pro Met Asn cct ccc aaa tat gac aag atc gag gat atg gcc atg atg act cat ctg Pro Pro Lys Tyr Asp Lys IIe Glu Asp Met Ala Met Met Thr His Leu cat gag cct gct gtg ctg tac aac ctc aaa gaa cgt tat gca gcc tgg His Glu Pro Ala Val Leu Tyr Asn Leu Lys Glu Arg Tyr Ala Ala Trp atg atc tac acc tat tca ggt ctc ttc tgt gtc act gtc aac ccc tac Met lle Tyr Thr Tyr Ser Gly Leu Phe Cys Val Thr Val Asn Pro Tyr aag tgg ctg cct gtg tat aag ccc gag gtg gtg aca gcc tac cga ggc Lys Trp Leu Pro Val Tyr Lys Pro Glu Val Val Thr Ala Tyr Arg Gly aaa aag cgc cag ggg gcc ccg ccc cac atc ttc tcc atc tct gac aac Lys Lys Arg Gln Gly Ala Pro Pro His IIe Phe Ser IIe Ser Asp Asn gcc tat cag ttc atg ctg act gac cga gag aat cag tca atc ctg atc

Ala Tyr Gin Phe Met Leu Thr Asp Arg Glu Asn Gin Ser ile Leu ile

		165	•		170			175				
		Ser		ggg Gly							69	93
				gca Ala 200							74	11
				cag Gln				-	He		. 78	9
				gcc Ala							. 83	7
-				ggt Gly							88	5
				gct Ala	•						93	3
				ctt Leu 280							98	1
				aag Lys						_	1029	9
				gat Asp					_		107	7

agt gtg gcc agc atc gat gat cag gaa gaa ctg atg gcc aca gat agt Ser Val Ala Ser lie Asp Asp Gin Glu Giu Leu Met Ala Thr Asp Ser gct att gat att ttg ggc ttt act aat gaa gaa aag gtc tcc att tac Ala lle Asp lle Leu Gly Phe Thr Asn Glu Glu Lys Val Ser ile Tyr aag ctc acg ggg gct gtg atg cat tat ggg aac cta aaa ttt aag caa Lys Leu Thr Gly Ala Val Met His Tyr Gly Asn Leu Lys Phe Lys Gln aag cag cgt gag gag caa gca gag cca gat ggc aca gaa gtt gct gac Lys Gln Arg Glu Glu Gln Ala Glu Pro Asp Gly Thr Glu Val Ala Asp aag gcg gcc tac ctc cag agt ctg aac tct gca gat ctg ctc aaa gct Lys Ala Ala Tyr Leu Gin Ser Leu Asn Ser Ala Asp Leu Leu Lys Ala ctc tgc tac ccc agg gtc aag gtc ggc aat gag tat gtc acc aaa ggc Leu Cys Tyr Pro Arg Val Lys Val Gly Asn Glu Tyr Val Thr Lys Gly cag act gta gaa cag gtg tcc aac gca gta ggt gct ctg gcc aaa gcc Gin Thr Val Giu Gin Val Ser Asn Ala Val Giy Ala Leu Ala Lys Ala gtc tac gag aag atg ttc ctg tgg atg gtt gcc cgc atc aac cag cag Val Tyr Giu Lys Met Phe Leu Trp Met Val Ala Arg !le Asn Gin Gin ctg gac acc aag cag ccc agg cag tac ttc atc ggg gtc ttg gac att Leu Asp Thr Lys Gin Pro Arg Gin Tyr Phe Ile Giy Val Leu Asp Ile

450	455	460	465
_		tc aac agc ctg gag c he Asn Ser Leu Glu G 475	
		aa cag ttt ttc aac c In GIn Phe Phe Asn H 490	
		ag aag gaa ggc atc g ys Lys Glu Gly lie G 05	•
		ct gcc tgc atc gag c la Ala Cys lle Glu Lo 525	•
_		ng gaa gag gag tgc at Bu Glu Glu Glu Cys Me 540	
		nc aag otg tat gac ca n Lys Leu Tyr Asp Gl 555	
Lys Ser Ala		c aag gtg gtc aaa gg o Lys Val Val Lys Gl 570	
		t gct ggt gtt gtg ga r Ala Gly Val Val As 5 59	p Tyr Asn lle
		g gac ccc ctg aat ga s Asp Pro Leu Asn Gl	-

gga ctg tac cag aag tot gca atg aaa act cta gct cag ctc ttc tct Gly Leu Tyr Gln Lys Ser Ala Met Lys Thr Leu Ala Gln Leu Phe Ser ggg gct caa act gct gaa gga gag gga gct ggc gga ggg gcc aag aaa Gly Ala Gin Thr Ala Glu Gly Glu Gly Ala Gly Gly Gly Ala Lys Lys ggt ggt aag aag agg ggc tot tot tto cag aca gtg tot gcc ott tto Gly Gly Lys Lys Gly Ser Ser Phe Gin Thr Vai Ser Ala Leu Phe aga gag aat ttg aac aag ctg atg acc aac ctc agg agt acc cat cct Arg Glu Asn Leu Asn Lys Leu Met Thr Asn Leu Arg Ser Thr His Pro 670 · cac ttt gtg agg tgt atc atc ccc aat gag aca aaa act cct ggt gcc His Phe Val Arg Cys IIe IIe Pro Asn Glu Thr Lys Thr Pro Gly Ala atg gag cat gag ctt gtc ctc cac cag ctg agg tgt aac ggt gtg ctg Met Glu His Glu Leu Val Leu His Gln Leu Arg Cys Asn Gly Val Leu gaa ggc atc cgc atc tgt agg aaa gga ttt cca agc aga atc ctt tat Glu Gly Ile Arg Ile Cys Arg Lys Gly Phe Pro Ser Arg Ile Leu Tyr gca gac ttc aaa cag aga tac aag gta tta aat gca agt gca atc cct Ala Asp Phe Lys Gin Arg Tyr Lys Val Leu Asn Ala Ser Ala Ile Pro gaa ggg caa tto att gat agc aag aag gcc tot gag aag ctc ott gca Glu Gly Gln Phe lle Asp Ser Lys Lys Ala Ser Glu Lys Leu Leu Ala

too atc gac att gac cac acc cag tat aaa ttt ggg cac acc aag gtc Ser lie Asp lie Asp His Thr Gln Tyr Lys Phe Gly His Thr Lys Val ttt ttc aaa gct ggt ctt ctg ggg ctc cta gag gag atg cga gat gac Phe Phe Lys Ala Gly Leu Leu Gly Leu Leu Glu Glu Met Arg Asp Asp aag ctg gcc cag ctg att acc cga acc cag gcc agg tgc aga ggg ttc Lys Leu Ala Gin Leu Ile Thr Arg Thr Gin Ala Arg Cys Arg Gly Phe ttg gca aga gtg gag tac cag agg atg gtg gag aga agg gag gcc atc Leu Ala Arg Val Glu Tyr Gln Arg Met Val Glu Arg Arg Glu Ala lie ttc tgt atc cag tac aat atc aga tcc ttc atg aat gtc aag cac tgg Phe Cys lie Gin Tyr Asn lie Arg Ser Phe Met Asn Val Lys His Trp ccc tgg atg aaa ctc ttc ttc aag atc aag cct ctg ttg aag agt gca Pro Trp Met Lys Leu Phe Phe Lys !le Lys Pro Leu Leu Lys Ser Ala gaa act gag aag gag atg gcc acc atg aag gaa gaa ttt cag aaa att Glu Thr Glu Lys Glu Met Ala Thr Met Lys Glu Glu Phe Gln Lys Ile aaa gac gaa ctt gcc aag tca gag gca aaa agg aag gaa ctg gaa gaa Lys Asp Glu Leu Ala Lys Ser Glu Ala Lys Arg Lys Glu Leu Glu Glu aag atg gtg acg ctg ttg aaa gaa aaa aat gac ttg cag ctc caa gtt Lys Met Val Thr Leu Leu Lys Glu Lys Asn Asp Leu Gln Val

cag gct gaa gcc gaa ggc ttg gct gat gca gag gaa agg tgt gac cag Gin Ala Giu Ala Giu Giy Leu Ala Asp Ala Giu Giu Arg Cys Asp Gin cta atc aaa acc aaa atc cag cta gaa gcc aaa atc aaa gag gtg act Leu lie Lys Thr Lys ile Gin Leu Glu Ala Lys lie Lys Glu Val Thr gag aga gct gag gat gag gaa gag atc aat gct gag ctg aca gcc aag Glu Arg Ala Glu Asp Glu Glu Glu Ile Asn Ala Glu Leu Thr Ala Lys . 945 aag agg aaa ctg gag gat gaa tgt tca gaa ctc aag aaa gac att gat Lys Arg Lys Leu Glu Asp Glu Cys Ser Glu Leu Lys Lys Asp ile Asp gac ctt gag ctg aca ctg gcc aag gtt gag aag gag aaa cat gcc aca Asp Leu Glu Leu Thr Leu Ala Lys Val Glu Lys Glu Lys His Ala Thr gaa aac aag gtg aaa aac ctc aca gaa gag atg gca ggt ctg gat gaa Glu Asn Lys Val Lys Asn Leu Thr Glu Glu Met Ala Gly Leu Asp Glu acc att gct aag ctg acc aag gag aag aag gct ctc cag gag gcc cac Thr lle Ala Lys Leu Thr Lys Glu Lys Lys Ala Leu Gln Glu Ala His cag cag acc ctg gat gac ctg cag gca gag gag gac aaa gtc aac Gin Gin Thr Leu Asp Asp Leu Gin Ala Giu Giu Asp Lys Val Asn acc ctg acc aaa gct aaa atc aaa ctt gaa caa caa gtg gat gat Thr Leu Thr Lys Ala Lys lie Lys Leu Glu Gin Gin Val Asp Asp

1025	1030	1035	
		aag aaa ctt cgc atg gac cta Lys Lys Leu Arg Met Asp Leu 1050	3276
		ggt gac ttg aag ttg gcc caa Gly Asp Leu Lys Leu Ala Gln 1065	3321
		gag aaa cag caa ctt gat gaa Glu Lys Gln Gln Leu Asp Glu 1080	. 3366
		atc agc aat ctg caa agc aag lle Ser Asn Leu Gin Ser Lys 1095	. 3411 . →
•		att caa ttg cag aag aaa att lle Gin Leu Gin Lys Lys lie 1110	3456
	caa goc ogo att gag g Gin Ala Arg lie Glu G 1120	gag ctg gag gag gaa atc gag Blu Leu Glu Glu Glu Ile Glu 1125	3501
		ca gag aag cag cgc tct gac la Glu Lys Gln Arg Ser Asp 1140	3546
		gc gag agg ctg gaa gaa gcc er Glu Arg Leu Glu Glu Ala 1155	3591
		ag atg aac aag aag cgg gag Iu Met Asn Lys Lys Arg Glu	3636

1160	110	65	1170	
		Arg Arg Asp L	etg gag gag gcc acc cta eu Glu Glu Ala Thr Leu 1185	3681
		Ala Thr Leu A	ngg aag aag cat gca gat nrg Lys Lys His Ala Asp 1200	3726
		Glu Gln ile A	ac aac ctg cag cga gtg sp Asn Leu Gin Arg Vai 1215	3771
		Glu Lys Ser G	ag atg aag atg gag att Iu Met Lys Met Glu Ile 1230	
-		Val Glu Thr Va	tc tcc aaa gcc aag gga al Ser Lys Ala Lys Gly 1245	3861
		Arg Thr Leu Gl	ng gac caa ctg agt gaa u Asp Gin Leu Ser Giu 1260	3906
		Glu Gln Gln Ar	g ctg atc aat gac ctg g Leu lie Asn Asp Leu 1275	3951
		Leu Gin Thr Gi	a tct ggt gag ttt tca u Ser Gly Glu Phe Ser 1290	3996
	t gat gaa aag	gaa gct ctg gtg	g tot cag tta tca aga I Ser Gin Leu Ser Arg	4041

1295			130	0 .		1305	5					
				Gin			Leu			g caa g Gln		4086
				Ala		ctg Leu 1335	Ala			ctg Leu		4131
				Cys		cgg Arg 1350	Glu			gag Glu	÷	4176
				Ala		aga Arg 1365	Ala		•		١.	. <mark>4221</mark> ,
				Gin		aaa Lys 1380					·	4266
				Glu		gcc Ala 1395						4311
gcc ca Ala G 1400						gta Val 1410		gct Ala				4356
						cag GIn 1425						4401
						agg Arg						4446

1430)				1435	5				1440)					
tgt Cys					aaa Lys							_		_		4491
1445					1450)				1455	;					
gca	gaa	tgg	aaa	cag	aaa	tgt	gag	gaa	acg	cat	gct	gag	ctt	gag		4536
Ala		Trp	Lys	Gin	Lys		Glu	Glu	Thr			Glu	Leu	Glu		
1460					1465					1470	•					
gcc		•			gcc										:	4581
Ala 1475		Gln	Lys	Glu	Ala 1480		Ser	Leu	Gly	Thr 1485		Leu	Phe	Lys		
										1400				•		
ata 					gag											4626
ile 1490	Lys	Asn	Ala	Tyr	GI u 1495		Ser	Leu	Asp	GIn 1500		Glu	Thr	Leu	1	•
					1400					1000						
aag					aac									_		4671
Lys 1505	Arg	Glu	Asn	Lys	Asn 1510		Gin	Gin	Glu	lle 1515	Ser	Asp	Leu	Thr		
.000					1010					1010						
gaa					gga							ctg	gag	aaa		4716
Glu 1520	Gin	He	Ala	Glu	Gly 1525	Gly	Lys	Arg	He	His 1530	Glu	Leu	Glu	Lys		
					1020					1000						
					gaa							cag				4761
11e	Lys	Lys	GIN	Val	Glu 1540	Gin	Glu	Lys	Cys	Glu 1545	Leu	Gin	Ala	Ala		
.000					10-10					1040						
					gca											4806
Leu 1550	Glu	Glu	Ala	Glu	Ala 1555	Ser	Leu	Giu	His	Glu 1560	Glu	Gly	Lys	He		
1000					1000					1000			•			
					gag									_		4851
Leu	Arg	He	GIn	Leu	Glu	Leu	Asn	Gln	Val	Lys	Ser	Glu	Vai	Asp		

1565	1570	1575	
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		cc atg cag agc acg ctg gat gct er Met Gin Ser Thr Leu Asp Ala 1605	
		cc att agg ctc aag aag aag atg Ia lle Arg Leu Lys Lys Lys Met 1620	•
		aa atc cag ctg aac cat gcc aac Iu lle Gin Leu Asn His Ala Asn 1635	•
		gg aac tac agg aac acc caa ggc rg Asn Tyr Arg Asn Thr Gin Giy 1650	5076
		s Leu Asp Asp Ala Leu Arg Ser 1665	5121
		g gcc atg gtg gag cgc aga gcc u Ala Met Val Glu Arg Arg Ala 1680	5166
		g gag ctg cgg gcc act ctg gaa u Glu Leu Arg Ala Thr Leu Glu 1695	5211
		c gca gaa cag gag ctc ctg gat e Ala Glu Gin Giu Leu Leu Asp	5256

1700	1705	1710
man and man and a		
		acc cag aac acc agc ctg 5301
1715		Thr Gin Asn Thr Ser Leu
1715	1720	1725
atc aac acc aag a	ag aag ctg gag aca g	gat att too caa atg caa 5346
lle Asn Thr Lys L	ys Lys Leu Glu Thr A	Asp lie Ser Gin Met Gin
1730	1735	1740
	· .	·
		cc cgc aat gca gaa gaa 5391
Gly Glu Met Glu A	spille Leu Gin Giu A	la Arg Asn Ala Glu Glu .
1745	1750	1755
		cc atg atg gct gag gag .5436
	la lle Thr Asp Ala A	la Met Met Ala Glu Glu 🕠 🔻
1760	1765	1770
		ac ctg gag cgg atg aag 5481
		is Leu Glu Arg Met Lys
1775	1780	1785
000 000 000 000		
		tg cag ctc cgt ctg gat 5526
Lys Asn Met Glu Gl 1790		eu Gin Leu Arg Leu Asp
1790	1795	1800
gag gct gag cag ct	g gcc ctg aag ggt gg	gg aag aag cag atc cag 5571
		gg aag aag cag atc cag 5571 ly Lys Lys Gin lie Gin
1805	1810	1815
		10.0
aaa ctg gag gcc ag	g gta cgg gag ctg ga	na gga gag gtt gag agt 5616
		u Gly Glu Val Glu Ser
1820	1825	1830
gag caa aag cgt aa	t got gag got gto aa	a ggt ctg cgc aaa cat 5661
Glu Gln Lys Arg Ası	n Ala Giu Ala Val Lys	s Gly Leu Arg Lys His

1835		1840	1845	
			tac cag acg gaa gaa gat aga Tyr Gin Thr Giu Giu Asp Arg 1860	
			ttg gta gat aaa ctt cag gca Leu Val Asp Lys Leu Gin Ala 1875	
			gct gag gag gct gag gaa caa Ala Glu Glu Ala Glu Glu Gln 1890	5796
			cgc aag ctc cag cat gag ctg Arg Lys Leu Gin His Glu Leu 1905	
			att gct gag tcc cag gtg aac lle Ala Glu Ser Gln Val Asn 1920	5886
			gtt cac aca aaa gtc ata agt Val His Thr Lys Val IIe Ser 1935	5931
gaa gag Glu Glu 1940	tga tcatgtco	etg atgccatgga	atgactgaag acaggcacaa	5980
aatgtgaca	t ctttggtcat	ttccctctgt aa	attattgtg tattotaccc tgttgcaaa	ıg 6040
gaaataaago	c atagggtagt	ttgcaaacaa aa	aaaaaaaa aaaaa	6085

<211> 1941

<212> PRT

<213> Homo sapiens

<400> 2

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35 40 45

Lys Gly Thr lie Gln Ser Arg Glu Gly Gly Lys Val Thr Val Lys Thr 50 55 60

Glu Gly Gly Ala Thr Leu Thr Val Lys Asp Asp Gln Val Phe Pro Met 70 75 80

Asn Pro Pro Lys Tyr Asp Lys Ile Glu Asp Met Ala Met Met Thr His
85 90 95

Leu His Glu Pro Ala Val Leu Tyr Asn Leu Lys Glu Arg Tyr Ala Ala 100 105 110

Trp Met IIe Tyr Thr Tyr Ser Gly Leu Phe Cys Val Thr Val Asn Pro
115 120 125

Tyr Lys Trp Leu Pro Vai Tyr Lys Pro Glu Vai Vai Thr Ala Tyr Arg 130 135 140

Gly Lys Lys Arg Gln Gly Ala Pro Pro His IIe Phe Ser IIe Ser Asp 145 150 155 160

Asn Ala Tyr Gin Phe Met Leu Thr Asp Arg Giu Asn Gin Ser ile Leu

165 170 175

11e Thr Gly Glu Ser Gly Ala Gly Lys Thr Val Asn Thr Lys Arg Val 180 185 190

lle Gln Tyr Phe Ala Thr lle Ala Val Thr Gly Glu Lys Lys Glu 195 200 205

Glu lle Thr Ser Gly Lys lle Gln Gly Thr Leu Glu Asp Gln lle lle 210 215 220

Ser Ala Asn Pro Leu Leu Glu Ala Phe Gly Asn Ala Lys Thr Val Arg 225 230 235 240

Asn Asp Asn Ser Ser Arg Phe Gly Lys Phe IIe Arg IIe His Phe Gly
245 250 255

Thr Thr Gly Lys Leu Ala Ser Ala Asp IIe Glu Thr Tyr Leu Leu Glu 260 265 270

Lys Ser Arg Val Val Phe Gln Leu Lys Ala Glu Arg Ser Tyr His Ile 275 280 285

Phe Tyr Gin lie Thr Ser Asn Lys Lys Pro Giu Leu lie Giu Met Leu 290 295 300

Leu IIe Thr Thr Asn Pro Tyr Asp Tyr Pro Phe Val Ser Gln Gly Glu 305 310 315 320

lie Ser Val Ala Ser Ile Asp Asp Gin Giu Giu Leu Met Ala Thr Asp 325 330 335

Ser Ala IIe Asp IIe Leu Gly Phe Thr Asn Glu Glu Lys Val Ser IIe 340 345 350

Tyr Lys Leu Thr Gly Ala Val Met His Tyr Gly Asn Leu Lys Phe Lys 355 360 365

Gin Lys Gin Arg Giu Giu Gin Ala Giu Pro Asp Giy Thr Giu Val Ala 370 375 380

Asp Lys Ala Ala Tyr Leu Gln Ser Leu Asn Ser Ala Asp Leu Leu Lys 385 390 395 400

Ala Leu Cys Tyr Pro Arg Val Lys Val Gly Asn Glu Tyr Val Thr Lys
405 410 415

Gly Gln Thr Val Glu Gln Val Ser Asn Ala Val Gly Ala Leu Ala Lys 420 425 430

Ala Val Tyr Glu Lys Met Phe Leu Trp Met Val Ala Arg ile Asn Gln 435 440 445

Gin Leu Asp Thr Lys Gin Pro Arg Gin Tyr Phe ile Gly Val Leu Asp 450 455 460

Ile Ala Gly Phe Glu Ile Phe Asp Phe Asp Ser Leu Glu Gln Leu Cys465470475480

Ile Asn Phe Thr Asn Glu Lys Leu Gln Gln Phe Phe Asn His His Met 485 490 495

Phe Val Leu Glu Glu Glu Glu Tyr Lys Lys Glu Gly lie Glu Trp Thr
500 505 510

Phe IIe Asp Phe Gly Met Asp Leu Ala Ala Cys IIe Glu Leu IIe Glu 515 520 525

Lys Pro Met Gly lie Phe Ser ile Leu Glu Glu Glu Cys Met Phe Pro 530 535 540

Lys Ala Thr Asp Thr Ser Phe Lys Asn Lys Leu Tyr Asp Gln His Leu 545 550 555 560

Gly Lys Ser Ala Asn Phe Gln Lys Pro Lys Val Val Lys Gly Lys Ala 565 570 575

Glu Ala His Phe Ala Leu Ile His Tyr Ala Gly Val Val Asp Tyr Asn 580 585 590

lie Thr Gly Trp Leu Glu Lys Asn Lys Asp Pro Leu Asn Glu Thr Val 595 600 605

Val Gly Leu Tyr Gln Lys Ser Ala Met Lys Thr Leu Ala Gln Leu Phe 610 615 620

, . . .

Ser Gly Ala Gln Thr Ala Glu Gly Glu Gly Ala Gly Gly Gly Ala Lys 625 630 635 640

Lys Gly Gly Lys Lys Gly Ser Ser Phe Gln Thr Val Ser Ala Leu 645 650 655

Phe Arg Glu Asn Leu Asn Lys Leu Met Thr Asn Leu Arg Ser Thr His 660 665 670

Pro His Phe Val Arg Cys lle lle Pro Asn Glu Thr Lys Thr Pro Gly 675 680 685

Ala Met Glu His Glu Leu Val Leu His Gln Leu Arg Cys Asn Gly Val 690 695 700 Leu Glu Gly Ile Arg Ile Cys Arg Lys Gly Phe Pro Ser Arg Ile Leu 705 710 715 720

Tyr Ala Asp Phe Lys Gln Arg Tyr Lys Val Leu Asn Ala Ser Ala ile 725 730 735

Pro Glu Gly Gin Phe Ile Asp Ser Lys Lys Ala Ser Glu Lys Leu Leu . 740 745 750

Ala Ser Ile Asp Ile Asp His Thr Gln Tyr Lys Phe Gly His Thr Lys
755 760 765

Val Phe Phe Lys Ala Gly Leu Leu Gly Leu Leu Glu Glu Met Arg Asp 770 775 780

Asp Lys Leu Ala Gln Leu IIe Thr Arg Thr Gln Ala Arg Cys Arg Gly 785 790 795 800

Phe Leu Ala Arg Val Glu Tyr Gln Arg Met Val Glu Arg Arg Glu Ala 805 810 815

ile Phe Cys Ile Gin Tyr Asn Ile Arg Ser Phe Met Asn Val Lys His 820 825 830

Trp Pro Trp Met Lys Leu Phe Phe Lys IIe Lys Pro Leu Lys Ser 835 840 845 Ala Glu Thr Glu Lys Glu Met Ala Thr Met Lys Glu Glu Phe Gin Lys 850 855 860

Ile Lys Asp Glu Leu Ala Lys Ser Glu Ala Lys Arg Lys Glu Leu Glu865870875880

Glu Lys Met Val Thr Leu Leu Lys Glu Lys Asn Asp Leu Gln Leu Gln . 885 890 895

Val Gln Ala Glu Ala Glu Gly Leu Ala Asp Ala Glu Glu Arg Cys Asp 900 905 910

Gin Leu IIe Lys Thr Lys IIe Gin Leu Glu Ala Lys IIe Lys Glu Val 915 920 925

Thr Glu Arg Ala Glu Asp Glu Glu Glu Ile Asn Ala Glu Leu Thr Ala 930 935 940

Lys Lys Arg Lys Leu Glu Asp Glu Cys Ser Glu Leu Lys Lys Asp I le 945 950 955 960

Asp Asp Leu Glu Leu Thr Leu Ala Lys Val Glu Lys Glu Lys His Ala 965 970 975

Thr Glu Asn Lys Vai Lys Asn Leu Thr Glu Glu Met Ala Gly Leu Asp 980 985 990 Glu Thr lle Ala Lys Leu Thr Lys Glu Lys Lys Ala Leu Gln Glu Ala 995 1000 1005

His Gln Gln Thr Leu Asp Asp Leu Gln Ala Glu Glu Asp Lys Val 1010 1015 1020

Asn Thr Leu Thr Lys Ala Lys lle Lys Leu Glu Gln Gln Val Asp 1025 1030 1035

Asp Leu Glu Gly Ser Leu Glu Gln Glu Lys Lys Leu Arg Met Asp 1040 1045 1050

1. 3

Leu Glu Arg Ala Lys Arg Lys Leu Glu Gly Asp Leu Lys Leu Ala 1055 1060 1065

Gin Glu Ser lie Met Asp lie Glu Asn Glu Lys Gin Gln Leu Asp 1070 1075 1080

Glu Lys Leu Lys Lys Glu Phe Glu lle Ser Asn Leu Gln Ser 1085 1090 1095

Lys Ile Giu Asp Giu Gin Ala Leu Giy Ile Gin Leu Gin Lys Lys 1100 1105 1110

lle Lys Glu Leu Gin Ala Arg lle Glu Glu Leu Glu Glu lle 1115 1120 1125

- Glu Ala Glu Arg Ala Ser Arg Ala Lys Ala Glu Lys Gln Arg Ser 1130 1135 1140
- Asp Leu Ser Arg Glu Leu Glu Glu IIe Ser Glu Arg Leu Glu Glu
 1145 1150 1155
- Ala Gly Gly Ala Thr Ser Ala Gln lle Glu Met Asn Lys Lys Arg 1160 1165 1170
- Glu Ala Glu Phe Gln Lys Met Arg Arg Asp Leu Glu Glu Ala Thr 1175 1180 1185

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- Leu Gln His Glu Ala Thr Ala Ala Thr Leu Arg Lys Lys His Ala 1190 1195 1200
- Asp Ser Val Ala Glu Leu Gly Glu Gln IIe Asp Asn Leu Gln Arg 1205 1210 1215
- Val Lys Gin Lys Leu Glu Lys Glu Lys Ser Glu Met Lys Met Glu 1220 1225 1230
- lle Asp Asp Leu Ala Ser Asn Val Glu Thr Val Ser Lys Ala Lys 1235 1240 1245
- Gly Asn Leu Glu Lys Met Cys Arg Thr Leu Glu Asp Gln Leu Ser 1250 1255 1260

- Glu Leu Lys Ser Lys Glu Glu Glu Gln Gln Arg Leu ile Asn Asp 1265 1270 1275
- Leu Thr Ala Gin Arg Gly Arg Leu Gin Thr Glu Ser Gly Glu Phe 1280 1285 1290
- Ser Arg Gin Leu Asp Giu Lys Giu Ala Leu Val Ser Gin Leu Ser 1295 1300 1305
- Arg Gly Lys Gln Ala Phe Thr Gln Gln He Glu Glu Leu Lys Arg 1310 1315 1320
- Gin Leu Glu Glu IIe Lys Ala Lys Asn Ala Leu Ala His Ala 1325 1330 1335
- Leu Gln Ser Ser Arg His Asp Cys Asp Leu Leu Arg Glu Gln Tyr 1340 1345 1350
- Glu Glu Glu Gln Glu Ser Lys Ala Glu Leu Gln Arg Ala Leu Ser 1355 1360 1365
- Lys Ala Asn Thr Glu Vai Ala Gln Trp Arg Thr Lys Tyr Glu Thr 1370 1375 1380
- Asp Ala ile Gin Arg Thr Giu Giu Leu Giu Giu Ala Lys Lys Lys 1385 1390 1395

- Leu Ala Gin Arg Leu Gin Ala Ala Giu Giu His Val Giu Ala Val 1400 1405 1410
- Asn Ala Lys Cys Ala Ser Leu Glu Lys Thr Lys Gln Arg Leu Gln 1415 1420 1425
- Asn Glu Val Glu Asp Leu Met Leu Asp Val Glu Arg Thr Asn Ala 1430 1435 1440
- Ala Cys Ala Ala Leu Asp Lys Lys Gin Arg Asn Phe Asp Lys Ile 1445 1450 1455
- Leu Ala Glu Trp Lys Gln Lys Cys Glu Glu Thr His Ala Glu Leu 1460 1465 1470
- Glu Ala Ser Gln Lys Glu Ala Arg Ser Leu Gly Thr Glu Leu Phe 1475 1480 1485
- Lys Ile Lys Asn Ala Tyr Glu Glu Ser Leu Asp Gln Leu Glu Thr 1490 1495 1500
- Leu Lys Arg Glu Asn Lys Asn Leu Gin Gin Glu IIe Ser Asp Leu 1505 1510 1515
- Thr Glu Gin He Ala Glu Gly Gly Lys Arg He His Glu Leu Glu 1520 1525 1530

- Lys lie Lys Lys Gin Val Glu Gin Giu Lys Cys Giu Leu Gin Ala 1535 1540 1545
- Ala Leu Glu Glu Ala Glu Ala Ser Leu Glu His Glu Glu Gly Lys 1550 1555 1560
- lie Leu Arg lie Gin Leu Glu Leu Asn Gin Val Lys Ser Glu Val 1565 1570 1575
- Asp Arg Lys lie Ala Giu Lys Asp Giu Giu lie Asp Gin Leu Lys 1580 1585 1590
- Arg Asn His IIe Arg IIe Val Glu Ser Met Gln Ser Thr Leu Asp 1595 1600 1605
- Ala Glu lle Arg Ser Arg Asn Asp Ala lle Arg Leu Lys Lys Lys 1610 1615 1620
- Met Glu Gly Asp Leu Asn Glu Met Glu lle Gln Leu Asn His Ala 1625 1630 1635
- Asn Arg Met Ala Ala Glu Ala Leu Arg Asn Tyr Arg Asn Thr Gln 1640 1645 1650
- Gly lle Leu Lys Asp Thr Gln lle His Leu Asp Asp Ala Leu Arg 1655 1660 1665

Ser Gin Giu Asp Leu Lys Giu Gin Leu Aia Met Vai Giu Arg Arg 1670 1675 1680

Ala Asn Leu Leu Gin Ala Giu Ile Giu Giu Leu Arg Ala Thr Leu 1685 1690 1695

Glu Gln Thr Glu Arg Ser Arg Lys I le Ala Glu Gln Glu Leu Leu 1700 1705 1710

Asp Ala Ser Glu Arg Val Gln Leu Leu His Thr Gln Asn Thr Ser 1715 1720 1725

Leu lle Asn Thr Lys Lys Leu Glu Thr Asp lle Ser Gln Met 1730 1735 1740

Gin Giy Giu Met Giu Asp ile Leu Gin Giu Ala Arg Asn Ala Giu 1745 1750 1755

Glu Lys Ala Lys Lys Ala IIe Thr Asp Ala Ala Met Met Ala Glu 1760 1765 1770

Glu Leu Lys Lys Glu Gln Asp Thr Ser Ala His Leu Glu Arg Met 1775 1780 1785

Lys Lys Asn Met Glu Gln Thr Val Lys Asp Leu Gln Leu Arg Leu 1790 1795 1800 Asp Glu Ala Glu Gln Leu Ala Leu Lys Gly Gly Lys Lys Gln lle 1805 1810 1815

Gin Lys Leu Giu Ala Arg Val Arg Giu Leu Giu Giy Giu Val Giu 1820 1825 1830

Ser Glu Gln Lys Arg Asn Ala Glu Ala Val Lys Gly Leu Arg Lys 1835 1840 1845

His Glu Arg Arg Val Lys Glu Leu Thr Tyr Gln Thr Glu Glu Asp 1850 1855 1860

Arg Lys Asn He Leu Arg Leu Gln Asp Leu Val Asp Lys Leu Gln 1865 1870 1875

Ala Lys Val Lys Ser Tyr Lys Arg Gin Ala Giu Giu Ala Giu Giu 1880 1885 1890

Gin Ser Asn Thr Asn Leu Ala Lys Phe Arg Lys Leu Gin His Giu 1895 1900 1905

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Asn Lys Leu Arg Val Lys Ser Arg Glu Val His Thr Lys Val IIe 1925 1930 1935 Ser Glu Glu 1940

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165

Ser Asp Ser Glu Met Ala IIe Phe Gly Glu Ala Ala Pro Phe Leu Arg

10

15

aag tot gaa aag gag cga att gaa got cag aac aag cot ttt gat goo 213 Lys Ser Glu Lys Glu Arg Ile Glu Ala Gln Asn Lys Pro Phe Asp Ala 20 25 30

aag aca tca gtc ttt gtg gtg gac cct aag gag tcc tac gtg aaa gca

261

Lys Thr Ser Val Phe Val Val Asp Pro Lys Glu Ser Tyr Val Lys Ala

35

40

45

50

ata gtg cag agc agg gaa ggg ggg aag gtg aca gcc aag acc gaa gct 309

ΙΙε	e Val	l Gir	n Sei	55	g Glu	Gly	Gl)	/ Lys	60	l Thr	- Ala	a Lys	s Thi	r Gli 65	ı Ala		
gga	gct	t act	gta	act	gtg	aaa	gaa	gac	caa	gto	tto	to	ate	g aad	cct		357
Gly	Ala	1 Thr	Val	Thr	· Val	Lys	Glu	ı Asp	Gir	Va I	Phe	e Ser	Met	: Asr	Pro		
			70					75					80				
ccc	aaa	tat	gac	aag	atc	gag	gac	atg	gcc	atg	ate	act	cac	cte	cat		405
Pro	Lys	Tyr	Asp	Lys	ile	Glu	Asp	Met	Ala	Met	Met	Thr	His	Leu	His	•	
		85					90					95					
gag	cct	gct	gtg	ctg	tat	aac	ctc	aaa	gag	cgt	tac	gca	gcc	tgg	atg	٠	453
Glu			Val	Leu	Tyr	Asn	Leu	Lys	Glu	Arg	Tyr	Ala	Ala	Trp	Met		
	100				•	105					110					·	
atc	tac	acc	tac	tcg	ggc	ctc	ttc	tgt	gtc	acc	gtc	aac	CCC	tac	aag	1. 9	501
He	Tyr	Thr	Tyr	Ser	Gly	Leu	Phe	Cys	Val	Thr	Val	Asn	Pro	Tyr	Lys	•	
.115					120					125					130		
tgg	ctg	ccg	gtg	tac	aac	cct	gag	gtg	gtg	aca	gcc	tac	cga	ggc	aaa		549
Trp	Leu	Pro	Val	Tyr	Asn	Pro	Glu	Val	Vai	Thr	Ala	Tyr	Arg	Gly	Lys		
				135					140					145			
aag	cgc	cag	gag	gcc	cca	ccc	cat	atc	ttc	tcc	atc	tct	gac	aat	gcc		597
Lys	Arg	GIn		Ala	Pro	Pro	His	lle	Phe	Ser	ile	Ser	Asp	Asn	Ala		
			150					155					160				
tat	cag	ttc	atg	cta	act	gat	cgt	gaa	aac	cag	tca	atc	ttg	att	act		645
Tyr	Gin		Met	Leu	Thr	Asp	Arg	Glu	Asn	Gln	Ser	He	Leu	lle	Thr		
		165					170					175			٠		
gga	gaa	tct	ggt	gca	ggg	aag	act	gtg	aac	acg	aag	cgt	gtc	atc	cag		693
		Ser	Gly	Ala	Gly	Lys	Thr	Val	Asn	Thr	Lys	Arg	Val	He	GIn		
	180					185					190						
tac	ttt	gca	aca	att	gca	gtt	act	gga	gag	aag	aaa	aaa	gag	gaa	cct	-	741

Tyr Phe Ala Thr Ile Ala Val Thr Gly Glu Lys Lys Glu Glu Pro	
195 200 205 210	0 .
gcc tct ggc aaa atg cag ggg acc ctt gaa gat caa atc atc agt gct	t 789
Ala Ser Gly Lys Met Gln Gly Thr Leu Glu Asp Gln He He Ser Ala	a .
215 220 225	
aac ccc cta ctg gaa gcc ttc ggc aat gcc aag acc gtg agg aat gac	837
Asn Pro Leu Leu Glu Ala Phe Gly Asn Ala Lys Thr Val Arg Asn Asp	
230 235 240	
	;
aac too tot ogo ttt ggt aaa tto ato agg ato cat ttt ggt goo aca Asn Ser Ser Arg Phe Gly Lys Phe Ile Arg Ile His Phe Gly Ala Thr	
245 250 255	
	_
ggc aaa ctg gct tct gca gat att gaa aca tat ctg cta gag aag tcc	
Gly Lys Leu Ala Ser Ala Asp Ile Glu Thr Tyr Leu Leu Glu Lys Ser	
260 265 270	
cga gtt act ttt cag cta aag gct gaa aga agc tac cac ata ttt tat	981
Arg Val Thr Phe Gin Leu Lys Ala Glu Arg Ser Tyr His lie Phe Tyr	301
275 280 285 290	
caa atc ctg tcc aat aag aaa cca gag ctc att gaa atg ctt ctg atc	1029
Gin ile Leu Ser Asn Lys Lys Pro Giu Leu ile Giu Met Leu Leu ile 295 300 305	
295 300 305	
acc acc aac cca tat gac ttc gca ttt gtc agc caa ggg gaa att act	1077
Thr Thr Asn Pro Tyr Asp Phe Ala Phe Val Ser Gin Gly Glu lie Thr	
310 315 320	
gtg CCC agc att gat gac ong goo gog otg otg otg	44.00
gtg ccc agc att gat gac cag gaa gag ctg atg gcc aca gat agt gct Val Pro Ser Ile Asp Asp Gin Giu Giu Leu Met Ala Thr Asp Ser Ala	1125
325 330 335	
gtg gac atc ctg ggt ttc act gct gat gaa aag gtg gcc att tac aag	1173

Val	As ₁		e Lei	u Gly	/ Phe	345		a As _i	p Gi	ı Ly:	s Va 35		a li	е Ту	r Lys		
															a aag		1221
		GIS	/ Ala	a Val	Met	His	Туг	Gly	/ Asr	n Met	t Ly:	s Pho	e Ly	s Gl	n Lys		
355					360	•				365	5				370		
									•						c aaa		1269
Gln	Arg	Glu	Glu	Gin	Ala	Glu	Pro	Asp	Gly	Thr	Glu	ı Vai	l Ala	a Ası	Lys		
				375				•	380)				38	5		
gct	gct	tat	ctg	aca	agt	ctg	aac	tct	gct	gac	cte	cto	; aaa	tc1	ctc	•	1317
Ala	Ala	Tyr	Leu	Thr	Ser	Leu	Asn	Ser	Ala	Asp	Leu	Leu	Lys	Ser	Leu		
			390	1				395					400)			
tgc	tat	ccc	aga	gtc	aag	gtc	ggc	aat	gag	ttc	gta	acc	aaa	ggo	cag	١.	∍1365
Cys	Tyr	Pro	Arg	Val	Lys	Val	Gly	Asn	Glu	Phe	Val	Thr	Lys	Gly	GIn	•	
•		405					410					415					
															atc		1413
Thr		Gin	GIn	Val	Tyr	Asn	Ala	Val	Gly	Ala	Leu	Ala	Lys	Ala	He		
	420					425		,			430						
				ttc													1461
	Glu	Lys	Met	Phe	Leu	Trp	Met	Val	Thr	Arg	He	Asn	Gin	Gln	Leu		
435					440					445					450		
gac																	1509
Asp	Thr	Lys	Gln	Pro	Arg	GIn	Tyr	Phe	He	Gly	Val	Leu	Asp	He	Ala		
				455					460					465			
ggc																	1557
Gly	Phe	Glu	He	Phe	Asp	Phe	Asn	Ser	Leu	Glu	Gln	Leu	Cys	He	Asn		
			470					475 ·					480				
ttc a	acc	aac	gag	aaa	ctg	caa (cag	ttt	ttc	aac	cac	cac	atg	ttc	gtg		1605

Phe Thr Asn Glu Lys Leu Gln Gln Phe Phe Asn His His Met Phe Val 485 490 495	
ctg gag cag gaa gag tac aag aag gaa ggc atc gag tgg gag ttc att	1653
Leu Giu Gin Giu Giu Tyr Lys Lys Giu Giy lle Giu Trp Giu Phe lle	
500 505 510	
gac ttc ggg atg gac ctg gct gcc tgc atc gag ctc atc gag aag cct	1701
Asp Phe Gly Met Asp Leu Ala Ala Cys IIe Glu Leu IIe Glu Lys Pro	•
515 520 525 530	
atg ggc atc ttc tcc atc cta gaa gag gag tgc atg ttc ccc aag gca	1740
Met Gly lie Phe Ser lie Leu Glu Glu Glu Cys Met Phe Pro Lys Ala	1749
535 540 545	
aca gac acc tcc ttc aag aac aag ctg tat gaa caa cat ctt gga aaa	. → 1797
Thr Asp Thr Ser Phe Lys Asn Lys Leu Tyr Glu Gln His Leu Gly Lys	
550 555 560	
-	
too aac aac tto cag aag coc aag cot goc aaa ggc aag cot gag got	1845
Ser Asn Asn Phe Gin Lys Pro Lys Pro Ala Lys Gly Lys Pro Glu Ala	
565 570 575	
cac ttc tca ctg gtg cac tat gcc ggc acc gtg gac tac aac atc gcc	1000
His Phe Ser Leu Val His Tyr Ala Gly Thr Val Asp Tyr Asn Ile Ala	1893
580 585 590	
ggc tgg ctg gac aaa aac aag gac ccc ctg aat gag act gtg gtg ggg	1941
Gly Trp Leu Asp Lys Asn Lys Asp Pro Leu Asn Glu Thr Val Val Gly	
595 600 605 610	
ctg tac cag aag tot gca atg aag act ctg gct ttc ctc tct ggg	1989
Leu Tyr Gin Lys Ser Ala Met Lys Thr Leu Ala Phe Leu Phe Ser Gly	
615 620 625	
TC2 C22 act got got got got got got got got got	
gca caa act gct gaa gca gag ggt ggt ggt gga aag aaa ggt ggc aaa	2037

Ala Gin Thr Ala Giu Ala Giu Giy Giy Giy Lys Lys Giy Giy Lys 630 635 640	•
aag aag ggt tot tot tto oag aca gtg toa gct ott tto agg gag aat Lys Lys Gly Ser Ser Phe Gln Thr Val Ser Ala Leu Phe Arg Glu Asn 645 650 655	2085
ttg aat aag ctg atg acc aac ttg agg agc act cac ccc cac ttt gtg Leu Asn Lys Leu Met Thr Asn Leu Arg Ser Thr His Pro His Phe Val 660 665 670	2133
cgg tgc atc atc ccc aat gaa act aaa act cct ggt gcc atg gag cat Arg Cys IIe IIe Pro Asn Glu Thr Lys Thr Pro Gly Ala Met Glu His 675 680 685 690	2181
gag ctt gtc ctg cat cag ctg agg tgt aac ggt gtg ctg gaa ggc atc Glu Leu Val Leu His Gin Leu Arg Cys Asn Gly Val Leu Glu Gly Ile 695 700 705	-2229
cgc atc tgc agg aaa ggc ttc cca agc aga atc ctt tat gca gac ttc Arg lle Cys Arg Lys Gly Phe Pro Ser Arg lle Leu Tyr Ala Asp Phe 710 715 720	2277
aaa cag aga tac aag gtt cta aat gcg agt gct atc cca gag ggt cag Lys Gin Arg Tyr Lys Val Leu Asn Ala Ser Ala ile Pro Giu Giy Gin 725 730 735	2325
ttc att gac agc aag aag gct tct gag aaa ctt cta ggg tct att gaa Phe lle Asp Ser Lys Lys Ala Ser Glu Lys Leu Leu Gly Ser lle Glu 740 745 750	2373
att gac cac acc cag tac aaa ttc ggt cat acc aag gtt ttc ttc aaa lle Asp His Thr Gln Tyr Lys Phe Gly His Thr Lys Val Phe Phe Lys 755 760 765 770	2421
gct ggc ctg ctg gga act cta gaa gaa atg cga gat gaa aag cta gct	2469

Ala Gly Leu Leu Gly Thr Leu Glu Glu Met Arg Asp Glu Lys Leu Ala 775 780 785	-
caa ctc atc acg cgc act caa gcc ata tgc agg ggg ttc ctg atg aga Gin Leu lie Thr Arg Thr Gin Ala ile Cys Arg Gly Phe Leu Met Arg 790 795 800	2517
gtg gag ttc aga aag atg atg gag agg aga gag tcc atc ttc tgc att Val Glu Phe Arg Lys Met Met Glu Arg Arg Glu Ser ile Phe Cys ile 805 810 815	2565
cag tac aac atc cgt gct ttc atg aat gtg aag cac tgg ccc tgg atg Gln Tyr Asn lle Arg Ala Phe Met Asn Val Lys His Trp Pro Trp Met 820 825 830	2613
aag ctg tat ttc aag atc aag ccc ctc ctc aag agt gca gag aca gag Lys Leu Tyr Phe Lys IIe Lys Pro Leu Leu Lys Ser Ala Glu Thr Glu 835 840 845 850	. →2661
aag gag atg gcc aac atg aag gaa gaa ttt gag aaa acc aaa gaa gag Lys Glu Met Ala Asn Met Lys Glu Glu Phe Glu Lys Thr Lys Glu Glu 855 860 865	2709
ctg gct aag aca gag gca aaa agg aaa gaa cta gaa gaa aag atg gtg Leu Ala Lys Thr Glu Ala Lys Arg Lys Glu Leu Glu Glu Lys Met Val 870 875 880	2757
acg cta atg caa gag aaa aat gac tta caa ctc caa gtt caa gct gaa Thr Leu Met Gin Giu Lys Asn Asp Leu Gin Leu Gin Vai Gin Ala Giu 885 890 895	2805
gca gat gcc ttg gct gat gca gag gaa aga tgt gat cag ttg att aaa Ala Asp Ala Leu Ala Asp Ala Glu Glu Arg Cys Asp Gin Leu Ile Lys 900 905 910	2853
acc aaa atc caa ctt gag gcc aaa atc aaa gag gta act gaa aga gct	2901

Thr Lys ile Gin Leu Giu Ala Lys ile Lys Giu Vai Thr Giu Arg Aia	
915 920 925 930	
gag gat gag gaa gag atc aat gct gag ctg aca gcc aag aag agg aaa	2949
Glu Asp Glu Glu Ile Asn Ala Glu Leu Thr Ala Lys Lys Arg Lys	
935 940 945	
ctg gag gat gaa tgt tca gag ctc aag aaa gac att gat gac ctt gag	2997
Leu Glu Asp Glu Cys Ser Glu Leu Lys Lys Asp Ile Asp Asp Leu Glu 950 955 960	
950 955 960	
ctg aca ctg gcc aag gtt gag aag gag aaa cat gcc aca gag aac aag	3045
Leu Thr Leu Ala Lys Val Glu Lys Glu Lys His Ala Thr Glu Asn Lys	
965 970 975	
gtg aaa aac ctc aca gaa gag atg gca ggt ctg gat gaa acc att gct	8093
Val Lys Asn Leu Thr Glu Glu Met Ala Gly Leu Asp Glu Thr lle Ala 980 985 990	
980 985 990	
aag ctg acc aag gag aag aag gct ctc cag gag gcc cac cag cag	3138
Lys Leu Thr Lys Glu Lys Lys Ala Leu Gln Glu Ala His Gln Gln	
995 1000 1005	
acc ctg gat gac ctg cag atg gag gag gac aaa gtc aac acc ctg Thr Leu Asp Asp Leu Gin Met Glu Asp Lys Val Asp Tha Lau	3183
1010	
1010 1015 1020	
acc aaa gct aaa acc aag cta gaa cag caa gtg gac gat ctt gaa	3228
or and looking out out out gig gat gat til gad	3228
	3228
Thr Lys Ala Lys Thr Lys Leu Glu Gin Gin Vai Asp Asp Leu Glu 1025 1030 1035	3228
Thr Lys Ala Lys Thr Lys Leu Glu Gln Gln Val Asp Asp Leu Glu 1025 1030 1035 gga tct ctg gaa caa gaa aag aaa ctt tgc atg gac tta gaa aga	3228 3273
Thr Lys Ala Lys Thr Lys Leu Glu Gln Gln Val Asp Asp Leu Glu 1025 1030 1035 gga tct ctg gaa caa gaa aag aaa ctt tgc atg gac tta gaa aga Gly Ser Leu Glu Gln Glu Lys Lys Leu Cys Met Asp Leu Glu Arg	
Thr Lys Ala Lys Thr Lys Leu Glu Gln Gln Val Asp Asp Leu Glu 1025 1030 1035 gga tct ctg gaa caa gaa aag aaa ctt tgc atg gac tta gaa aga	
Thr Lys Ala Lys Thr Lys Leu Glu Gln Gln Val Asp Asp Leu Glu 1025 1030 1035 gga tct ctg gaa caa gaa aag aaa ctt tgc atg gac tta gaa aga Gly Ser Leu Glu Gln Glu Lys Lys Leu Cys Met Asp Leu Glu Arg	

Ala Lys Arg Lys Leu Glu Gly Asp Leu Lys Leu Ala Gln Glu Ser 1055 1060 1065	
aca atg gat aca gaa aat gac aaa cag caa ctt aat gag aaa ctc Thr Met Asp Thr Glu Asn Asp Lys Gln Gln Leu Asn Glu Lys Leu 1070 1075 1080	3363
aaa aag aaa gag ttt gaa atg agc aat ctg caa ggc aag att gaa Lys Lys Lys Glu Phe Glu Met Ser Asn Leu Gln Gly Lys Ile Glu 1085 1090 1095	3408
gat gaa caa gcc ctt gca atg cag cta caa aag aag atc aaa gaa Asp Glu Gin Ala Leu Ala Met Gin Leu Gin Lys Lys ile Lys Giu 1100 1105 1110	3453
tta cag gcc cgc att gag gag ctg gag gag gaa atc gag gca gag Leu Gin Ala Arg lie Giu Giu Leu Giu Giu Giu Ile Giu Ala Giu 1115 1120 1125	. → 3498
cgg gcc tcc cgg gcc aaa gca gaa aag cag cgc tct gac ctc tcc Arg Ala Ser Arg Ala Lys Ala Glu Lys Gln Arg Ser Asp Leu Ser 1130 1135 1140	3543
cgg gag ctg gag gag atc agt gag agg ctg gaa gaa gcc ggt ggg Arg Glu Leu Glu Glu IIe Ser Glu Arg Leu Glu Glu Ala Giy Gly 1145 1150 1155	3588
gcc act toa gcc cag att gag ttg aac aag aag cgg gag gct gag Ala Thr Ser Ala Gin iie Giu Leu Asn Lys Lys Arg Giu Ala Giu 1160 1165 1170	3633
ttc cag aaa atg cgc agg gac ctg gaa gag tcc acc ctg cag cac Phe Gln Lys Met Arg Arg Asp Leu Glu Glu Ser Thr Leu Gln His 1175 1180 1185	3678
gaa gcc acg gca gct gct ctt cgg aag aag cac gca gat agt gtg	3723

Glu Ala Thr Ala Ala Ala Leu Arg Lys Lys His Ala Asp Ser Va 1190 1195 1200	
gct gag ctt ggg aag cag atc gac agc ctt cag cgg gtc aag cag Ala Glu Leu Gly Lys Gln Ile Asp Ser Leu Gln Arg Val Lys Gli 1205 1210 1215	
aag ctg gag aag gaa aag agt gag ctg aag atg gag atc aat gad Lys Leu Glu Lys Glu Lys Ser Glu Leu Lys Met Glu lie Asn Asp 1220 1225 1230	
ctt gct agt aac atg gag act gtc tcc aaa gcc aag gca aac ttt Leu Ala Ser Asn Met Glu Thr Val Ser Lys Ala Lys Ala Asn. Phe 1235 1240 1245	
gag aaa atg tgc cgc acc cta gag gac cag ctt agt gaa ata aaa Glu Lys Met Cys Arg Thr Leu Glu Asp Gln Leu Ser Glu Ile Lys 1250 1255 1260	
aca aag gaa gaa gag cag caa cgc tta ata aat gag ttg tca gcc Thr Lys Glu Glu Glu Gln Gln Arg Leu lle Asn Glu Leu Ser Ala 1265 1270 1275	
cag aag gca cgt tta cac aca gaa tca ggt gag ttt tca cga cag Gln Lys Ala Arg Leu His Thr Glu Ser Gly Glu Phe Ser Arg Gln 1280 1285 1290	3993
cta gat gaa aaa gat gct atg gtt tct cag cta tcc cga ggc aaa Leu Asp Glu Lys Asp Ala Met Val Ser Gln Leu Ser Arg Gly Lys 1295 1300 1305	4038
caa gca ttt aca caa cag att gaa gaa tta aag agg cag cta gaa Gin Ala Phe Thr Gin Gin lie Giu Giu Leu Lys Arg Gin Leu Giu 1310 1315 1320	4083
gag gag act aag gcc aag agc act ctg gcc cat gcc ctg cag tca	4128

Glu Glu Thr Lys Ala Lys Ser Thr Leu Ala His Ala Leu Gln Ser 1325 1330 1335	
gcc cgc cat gac tgt gac ctg ctg cgg gaa cag tat gag gag gag Ala Arg His Asp Cys Asp Leu Leu Arg Glu Gln Tyr Glu Glu Glu 1340 1345 1350	4173
cag gaa gcc aag gct gag ctg cag agg gga atg tcc aag gcc aac Gin Giu Aia Lys Aia Giu Leu Gin Arg Giy Met Ser Lys Aia Asn 1355 1360 1365	4218
agt gag gtt gcc cag tgg agg acc aag tac gag acg gac gcc atc Ser Glu Val Ala Gin Trp Arg Thr Lys Tyr Glu Thr Asp Ala lie 1370 1375 1380	4263
cag cgc aca gag gag ctg gag gag gcc aag aag aag cta gcc cag Gin Arg Thr Giu Giu Leu Giu Giu Aia Lys Lys Lys Leu Aia Gin 1385 1390 1395	4308
cgt ctg cag gat gca gaa gaa cat gta gaa gct gtg aat tcc aaa Arg Leu Gin Asp Ala Giu Giu His Val Giu Ala Val Asn Ser Lys 1400 1405 1410	4353
tgt gct tct ctt gaa aag aca aag cag agg cta cag aat gaa gta Cys Ala Ser Leu Glu Lys Thr Lys Gln Arg Leu Gln Asn Glu Val 1415 1420 1425	4398
gag gac ctc atg att gat gtg gaa cga tct aat gct gcc tgc ata Glu Asp Leu Met IIe Asp Val Glu Arg Ser Asn Ala Ala Cys IIe 1430 1435 1440	4443
gct ctc gat aag aag caa aga aac ttt gac aag gtt ctg gca gaa Ala Leu Asp Lys Lys Gin Arg Asn Phe Asp Lys Val Leu Ala Giu 1445 1450 1455	4488
tgg aaa cag aag tat gag gaa act cag gct gaa ctt gag gcc tcc	4533

Trp	Ly	s GI	n Ly	s Ty	r Glu	GI	u Th	r GI	n Al	a Glu	Leu G	lu Al	a Ser	
1460					146					147				
														•
cag	aag	g ga	g to	g cg	t tct	ct	c ag	c ac	t ga	g ctg	ttc a	ag gt	g aag	4578
Gln											Phe Ly			.0.0
1475					148					148		,	,0	•
				•							_			
aat	gco	; ta	c ga	g ga	a tcc	ct	g gat	t ca	t cti	t gaa	act ct	a aa	g Cga	4623
Asn											Thr Le		_	
1490					149		•			1500				
											-			
gag	aat	aa	g aa	c tta	a caa	cag	g gag	g att	tct	gac	ctg ac	a gas	z caa	: 4668
Glu											Leu Th			1000
1505					1510					1515				•
att	gca	gag	g ggt	t gga	aag	cat	ato	cat	gaa:	ctg	gag aa	a gta	1 225	, ⊸4713
He											Glu Ly			, .4710
1520				_	1525					1530			-,0	
aaa	caa	ctt	gat	: cat	gag	aag	agt	gaa	cta	cag	act to	c cta	gag	4758
Lys											Thr Se			4700
1535					1540					1545			u.u	
gaa	gca	gag	gca	tct	ctt	gag	cat	gaa	gaa	ggc	aaa at	t ctt	CEC	4803
Glu											Lys 116		_	4000
1550					1555					1560			, u B	
att	caa	ctt	gag	cta	aat	cag	gtg	aaa	tct	gag	att gad	c cga	aaa	4848
											lle Asp			10.10
1565					1570			•		1575			_,	
att	gct	gaa	aaa	gat	gaa	gaa	ctc	gat	cag	cta	aag agg	aac	cat	4893
											Lys Arg			.500
1580			-	•	1585			- 1-		1590				
										. = 2 3				
ctc	aga	gtt	gtg	gag	tca	atg	cag	agt	aca	ctg	gat gct	gar	atc	4938
						_				-3	J 60L	0~0		7300

Leu	Arg	Val	Val	Glu	ı Ser	Met	Glr	ı Ser	- Thi	Leu	As	p Ala	a Gl	u lle	
1595	j				1600)				160	5				
															•
agg	ago	aga	ı aat	: gat	gct	cte	e age	ato	: 225	. aag	22	o at	a aə	g gga	4983
Arg														u Gly	7303
1610		VI E	, ASI	ı nop			Aig	, , , , ,	Lys			s we	וטו	u GIY	•
1010					1615)				1620)				
gat														c cag	5028
Asp	Leu	Asn	Glu	Met	Gļu	He	GIn	Leu	Asn	His	Ala	a Asr	ı Arı	g Gln	•
1625					1630)				1635	5				
				٠.							•				÷
gct	gct	gag	gca	cta	agg	aat	ctt	aga	aac	aca	caa	ı gga	ata	a ctg	5073
Ala	Ala	Glu	Ala	Leu	Arg	Asn	Leu	Arg	Asn	Thr	Gir	ı Giv	, 11e	e .Leu	
1640					1645					1650					
aag	gac	act	cag	cta	cat	+ + \(\sigma	orat	πat		ato	0.55			gat	5110
_															. 5118
	лор	1111	uin	Leu			ASP	ASP	АТА	ile		GIY	Gir	1 ASP	
1655					1660					1665)				
-		•		•											
gac										cgc					5163
Asp	Leu	Lys	Glu	Gln	Leu	Ala	Met	Val	Glu	Arg	Arg	Ala	Asn	Leu	
1670					1675					1680					
atg	cag	gct	gaa	gtt	gaa	gag	ctc	agg	gca	tcc	ctg	gaa	cgg	act	5208
										Ser					
1685					1690					1695			6	••••	
gag	202	aac	200	222	at m	400	~~	000	~~~	ctt					5050
															5253
Glu	AIG	ury	AIB	LyS		Ala	Giu	GIN	GIU		Leu	Asp	Ala	Ser	
1700	_				1705					1710					
gaa	cgt	gtg	caa	ctt	ctg	cac	act	cag	aac	acc	agc	ctg	atc	aac	5298
Glu	Arg	Val	Gln	Leu	Leu	His	Thr	Gln	Asn	Thr	Ser	Leu	He	Asn	
1715					1720					1725					
acc	aag	aag	aag	ctg	gaa	aca	gac	att	tcc	caa	atc	Cag	gga	gaø	5343
	_	_	_	_									004	0~0	5575

Thr - 1730		Lys	Lys	Leu	Glu 1735	Thr	Asp	lle	Ser	GIn 1740	lle	Gin	Gly	Glu	
atg Met 1745				-	cag Gin 1750								_	_	5388
aag Lys 1760					gat Asp 1765										5433 ·
aag Lys 1775					agc Ser 1780										5478
atg Met 1790		_			aag Lys 1795		_	_		_		_			, ∗5 523
gag Glu 1805					aag Lys 1810										5568
gag Glu 1820				-	gag Glu 1825		-								5613
					gct Ala 1840										5658
					act Thr 1855										5703
att	ctc	agg	ctg	cag	gac	ttg	gtg	gac	aaa	ttg	caa	acc	aaa	gtc	5748

He	Leu	Arg	Leu	Gln	Asp	Leu	Val	Asp	Lys	Leu	Gin	Thr	Lys	Val	
1865				•	1870					1875					
															•
aaa	gct	tac	aag	aga	caa	gct	gaa	gag	gct	gag	gaa	caa	tcc	aat	5793
Lys	Ala	Tyr	Lys	Arg	Gln	Ala	Glu	Glu	Ala	Glu	Glu	Gin	Ser	Asn	
1880					1885					1890					
gtc	aac	ctt	gcc	aag	ttc	cgc	aag	ctc	cag	cac	gag	ctg	gag	gag	5838
Val	Asn	Leu	Ala	Lys	Phe	Arg	Lys	Leu	Gln	His	Glu	Leu	Glu	Glu	•
1895					1900					1905					
gcc	gag	gaa	cgg	gct	gac	att	gct	gag	tcc	caa	gtc	aac	aag	ctg	5883
Ala	Glu	Glu	Arg	Ala	Asp	He	Ala	Glu	Ser	Gln	Val	Asn	Lys	Leu	
1910					1915					1920					
aga	gtg	aag	agt	cgg	gag	gtt	cac	aca	aaa	gtc	ata	agt	gaa	gag	5928
Arg	Val	Lys	Ser	Arg	Glu	Val	His	Thr	Lys	Val	He	Ser	Glu	Glu	
1925					1930					1935					
_															
taa t	tcat	tcta	a tg	aaag	aaaa	tgtg	acca	aa g	aaat	gcace	; aaa	tgtg	aag		5981
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<213≻ Homo sapiens

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Lys Ala IIe Val Gin Ser Arg Giu Giy Giy Lys Val Thr Ala Lys Thr 50 55 60

Glu Ala Gly Ala Thr Val Thr Val Lys Glu Asp Gln Val Phe Ser Met 65 70 75 . 80

Leu His Glu Pro Ala Val Leu Tyr Asn Leu Lys Glu Arg Tyr Ala Ala 100 105 110

Trp Met IIe Tyr Thr Tyr Ser Gly Leu Phe Cys Val Thr Val Asn Pro 115 120 125

Tyr Lys Trp Leu Pro Val Tyr Asn Pro Glu Val Val Thr Ala Tyr Arg 130 135 140

Gly Lys Lys Arg Gln Glu Ala Pro Pro His IIe Phe Ser IIe Ser Asp 145 150 155 160

Asn Ala Tyr Gin Phe Met Leu Thr Asp Arg Giu Asn Gin Ser lie Leu

. 165 170 175

Ile Thr Gly Glu Ser Gly Ala Gly Lys Thr Val Asn Thr Lys Arg Val 180 185 190

lie Gin Tyr Phe Ala Thr lie Ala Val Thr Gly Glu Lys Lys Glu
195 200 . 205

Glu Pro Ala Ser Gly Lys Met Gln Gly Thr Leu Glu Asp Gln Ile Ile 210 215 220 .

Ser Ala Asn Pro Leu Leu Glu Ala Phe Gly Asn Ala Lys Thr Val Arg 225 230 235 240

Asn Asp Asn Ser Ser Arg Phe Gly Lys Phe IIe Arg IIe His Phe Gly
245 250 255

Ala Thr Gly Lys Leu Ala Ser Ala Asp IIe Glu Thr Tyr Leu Leu Glu 260 265 270

Lys Ser Arg Val Thr Phe Gln Leu Lys Ala Glu Arg Ser Tyr His 11e 275 280 285

Phe Tyr Gin IIe Leu Ser Asn Lys Lys Pro Giu Leu IIe Giu Met Leu 290 295 300

Leu ile Thr Thr Asn Pro Tyr Asp Phe Ala Phe Val Ser Gin Gly Glu

305 310 315 320

lle Thr Val Pro Ser lle Asp Asp Gin Glu Glu Leu Met Ala Thr Asp 325 330 335

Ser Ala Val Asp Ile Leu Gly Phe Thr Ala Asp Glu Lys Val Ala Ile 340 345 350

Tyr Lys Leu Thr Gly Ala Val Met His Tyr Gly Asn Met Lys Phe Lys 355 360 365 .

Gin Lys Gin Arg Giu Giu Gin Ala Giu Pro Asp Giy Thr Giu Vai Ala ,... 370 375 380

Asp Lys Ala Ala Tyr Leu Thr Ser Leu Asn Ser Ala Asp Leu Leu Lys 385 390 395 400

Ser Leu Cys Tyr Pro Arg Val Lys Val Gly Asn Glu Phe Val Thr Lys 405 410 415

Gly Gln Thr Val Gln Gln Val Tyr Asn Ala Val Gly Ala Leu Ala Lys 420 425 430

Ala ile Tyr Giu Lys Met Phe Leu Trp Met Val Thr Arg ile Asn Gin
435
440
445

Gin Leu Asp Thr Lys Gin Pro Arg Gin Tyr Phe Ile Gly Val Leu Asp

450 455 460

ile Ala Gly Phe Glu lle Phe Asp Phe Asn Ser Leu Glu Gln Leu Cys 465 470 475 480

lle Asn Phe Thr Asn Glu Lys Leu Gln Gln Phe Phe Asn His His Met
485 490 495

Phe Val Leu Glu Glu Glu Glu Tyr Lys Lys Glu Gly Ile Glu Trp Glu
500 505 510 .

Phe lie Asp Phe Gly Met Asp Leu Ala Ala Cys lie Glu Leu lie Glu ... 515 520 525

Lys Pro Met Gly lie Phe Ser lie Leu Glu Glu Glu Cys Met Phe Pro 530 535 540

Lys Ala Thr Asp Thr Ser Phe Lys Asn Lys Leu Tyr Glu Gln His Leu 545 550 555 560

Gly Lys Ser Asn Asn Phe Gln Lys Pro Lys Pro Ala Lys Gly Lys Pro 565 570 575

Glu Ala His Phe Ser Leu Val His Tyr Ala Gly Thr Val Asp Tyr Asn 580 585 590

lle Ala Gly Trp Leu Asp Lys Asn Lys Asp Pro Leu Asn Glu Thr Val

595 600 605

Val Gly Leu Tyr Gln Lys Ser Ala Met Lys Thr Leu Ala Phe Leu Phe 610 615 620

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Phe Val Arg Cys IIe IIe Pro Asn Glu Thr Lys Thr Pro Gly Ala Met 675 680 685

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Gly lie Arg lie Cys Arg Lys Gly Phe Pro Ser Arg lie Leu Tyr Ala 705 710 715 720

Asp Phe Lys Gin Arg Tyr Lys Val Leu Asn Ala Ser Ala IIe Pro Giu 725 730 735

Gly Gln Phe Ile Asp Ser Lys Lys Ala Ser Glu Lys Leu Leu Gly Ser

740 745 750

Ile Glu lle Asp His Thr Gln Tyr Lys Phe Gly His Thr Lys Val Phe
755 760 765

Phe Lys Ala Gly Leu Leu Gly Thr Leu Glu Glu Met Arg Asp Glu Lys
770 780

Leu Ala Gin Leu IIe Thr Arg Thr Gin Ala IIe Cys Arg Gly Phe Leu 785 790 795 . 800

Met Arg Val Glu Phe Arg Lys Met Met Glu Arg Arg Glu Ser IIe Phe ,... 805 810 815

Cys lle Gln Tyr Asn lle Arg Ala Phe Met Asn Val Lys His Trp Pro 820 825 830

Trp Met Lys Leu Tyr Phe Lys IIe Lys Pro Leu Leu Lys Ser Ala Glu 835 840 845

Thr Glu Lys Glu Met Ala Asn Met Lys Glu Glu Phe Glu Lys Thr Lys 850 855 860

Glu Glu Leu Ala Lys Thr Glu Ala Lys Arg Lys Glu Leu Glu Glu Lys 865 870 875 880

Met Val Thr Leu Met Gin Glu Lys Asn Asp Leu Gin Leu Gin Val Gin

885 890 895

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Asn Lys Val Lys Asn Leu Thr Glu Glu Met Ala Gly Leu Asp Glu Thr 980 985 990

Ile Ala Lys Leu Thr Lys Glu Lys Lys Ala Leu Gln Glu Ala His Gln 995 1000 1005

Gin Thr Leu Asp Asp Leu Gin Met Giu Giu Asp Lys Val Asn Thr 1010 1015 1020 .

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1025 1030 1035

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Ser Thr Met Asp Thr Glu Asn Asp Lys Gln Gln Leu Asn Glu Lys 1070 1075 1080

Leu Lys Lys Glu Phe Glu Met Ser Asn Leu Gln Gly Lys Ile , 1085 1090 1095

.Glu Asp Giu Gin Ala Leu Ala Met Gin Leu Gin Lys Lys Ile Lys 1100 1105 1110

Glu Leu Gln Ala Arg Ile Glu Glu Leu Glu Glu Glu Ile Glu Ala 1115 1120 1125

Glu Arg Ala Ser Arg Ala Lys Ala Glu Lys Gin Arg Ser Asp Leu 1130 1135 1140

Ser Arg Glu Leu Glu Glu IIe Ser Glu Arg Leu Glu Glu Ala Gly 1145 1150 1155

Gly Ala Thr Ser Ala Gln IIe Glu Leu Asn Lys Lys Arg Glu Ala

1160 1165 1170

Glu Phe Gln Lys Met Arg Arg Asp Leu Glu Glu Ser Thr Leu Gln 1175 1180 1185

His Glu Ala Thr Ala Ala Ala Leu Arg Lys Lys His Ala Asp Ser 1190 1195 1200

Val Ala Glu Leu Gly Lys Gln lle Asp Ser Leu Gln Arg Val Lys 1205 1210 1215

Gin Lys Leu Giu Lys Giu Lys Ser Giu Leu Lys Met Giu ile Asn ... , 1220 1225 1230

Asp Leu Ala Ser Asn Met Giu Thr Val Ser Lys Ala Lys Ala Asn 1235 1240 1245

Phe Glu Lys Met Cys Arg Thr Leu Glu Asp Gln Leu Ser Glu IIe 1250 1255 1260

Lys Thr Lys Glu Glu Glu Gln Gln Arg Leu Ile Asn Glu Leu Ser 1265 1270 1275

Ala Gln Lys Ala Arg Leu His Thr Glu Ser Gly Glu Phe Ser Arg 1280 1285 1290

Gin Leu Asp Giu Lys Asp Ala Met Val Ser Gin Leu Ser Arg Gly

1295 1300 1305

Lys Gin Ala Phe Thr Gin Gin lie Giu Giu Leu Lys Arg Gin Leu 1310 1315 1320

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Ser Ala Arg His Asp Cys Asp Leu Leu Arg Glu Gln Tyr Glu Glu 1340 1345 1350 .

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Asn Ser Glu Val Ala Gin Trp Arg Thr Lys Tyr Glu Thr Asp Ala 1370 1375 1380

lle Gin Arg Thr Glu Glu Leu Glu Glu Ala Lys Lys Lys Leu Ala 1385 1390 1395

Gln Arg Leu Gln Asp Ala Glu Glu His Val Glu Ala Val Asn Ser 1400 1405 1410

Lys Cys Ala Ser Leu Glu Lys Thr Lys Gln Arg Leu Gln Asn Glu 1415 1420 1425

Val Glu Asp Leu Met ile Asp Val Glu Arg Ser Asn Ala Ala Cys

1430 1435 1440

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Glu Trp Lys Gln Lys Tyr Glu Glu Thr Gln Ala Glu Leu Glu Ala 1460 1465 1470

Ser Gin Lys Giu Ser Arg Ser Leu Ser Thr Giu Leu Phe Lys Val 1475 1480 1485

Lys Asn Ala Tyr Glu Glu Ser Leu Asp His Leu Glu Thr Leu Lys ...,
1490 1495 1500

Arg Glu Asn Lys Asn Leu Gin Gin Glu lie Ser Asp Leu Thr Glu 1505 1510 1515

Gin lie Ala Glu Gly Gly Lys His IIe His Glu Leu Glu Lys Val 1520 1525 1530

Lys Lys Gln Leu Asp His Glu Lys Ser Glu Leu Gln Thr Ser Leu 1535 1540 1545

Glu Glu Ala Glu Ala Ser Leu Glu His Glu Glu Gly Lys Ile Leu 1550 1560

Arg lie Gin Leu Glu Leu Asn Gin Val Lys Ser Giu lie Asp Arg

1565 1570 1575

Lys Ile Ala Giu Lys Asp Giu Giu Leu Asp Gin Leu Lys Arg Asn 1580 1585 1590

His Leu Arg Val Val Glu Ser Met Gln Ser Thr Leu Asp Ala Glu 1595 1600 1605

Ile Arg Ser Arg Asn Asp Ala Leu Arg Ile Lys Lys Met Glu 1610 1615 1620 .

Gly Asp Leu Asn Glu Met Glu IIe Gln Leu Asn His Ala Asn Arg 1625 1630 1635

Gin Ala Ala Giu Ala Leu Arg Asn Leu Arg Asn Thr Gin Giy ile 1640 1645 1650

Leu Lys Asp Thr Gln Leu His Leu Asp Asp Ala lle Arg Gly Gln 1655 1660 1665

Asp Asp Leu Lys Glu Gln Leu Ala Met Val Glu Arg Arg Ala Asn 1670 1675 1680

Leu Met Gin Ala Giu Vai Giu Giu Leu Arg Ala Ser Leu Giu Arg 1685 1690 1695

Thr Glu Arg Gly Arg Lys Met Ala Glu Gln Glu Leu Leu Asp Ala

1700 1705 1710

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Asn Thr Lys Lys Leu Glu Thr Asp lie Ser Gin lie Gin Gly
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Lys Lys Glu Gln Asp Thr Ser Ala His Leu Glu Arg Met Lys Lys 1775 1780 1785

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Gln Lys His Asn Val Glu Ala Val Lys Gly Leu Arg Lys His Glu

1835 1840 1845

Arg Arg Val Lys Glu Leu Thr Tyr Gln Thr Glu Glu Asp Arg Lys 1850 1855 1860

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ggt caa act gtg cag cag gtg tac aat gca gtg ggt gct ctg gcc aaa Gly Gln Thr Val Gln Gln Val Tyr Asn Ala Val Gly Ala Leu Ala Lys gct gtc tac gat aag atg ttc ttg tgg atg gtc acc cgc atc aac cag Ala Val Tyr Asp Lys Met Phe Leu Trp Met Val Thr Arg Ile Asn Gin cag ctg gac acc aag cag ccc agg cag tac ttc att ggg gtc ttg gac Gin Leu Asp Thr Lys Gin Pro Arg Gin Tyr Phe lie Gly Val Leu Asp att gct ggc ttt gag atc ttt gat ttc aac agc ctg gag cag ctg tgc lle Ala Gly Phe Glu lle Phe Asp Phe Asn Ser Leu Glu Gln Leu Cys atc aac ttc acc aat gag aaa ctg caa cag ttt ttc aac cac cac atg lle Asn Phe Thr Asn Glu Lys Leu Gln Gln Phe Phe Asn His His Met ttc gtg ctg gag cag gag gag tac aag aag gaa ggc att gag tgg acg Phe Val Leu Glu Glu Glu Glu Tyr Lys Lys Glu Gly Ile Glu Trp Thr ttc att gac ttt ggg atg gac ctg gct gcc tgc atc gag ctc atc gag Phe lie Asp Phe Gly Met Asp Leu Ala Ala Cys lle Glu Leu lle Glu aag cct atg ggc atc ttc tcc atc ctg gaa gag gag tgc atg ttc ccc Lys Pro Met Gly Ile Phe Ser Ile Leu Glu Glu Glu Cys Met Phe Pro

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Lys Ala Thr Asp Thr Ser Phe Lys Asn Lys Leu Tyr Glu Gln His Leu

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	ı Tyr Glin Lys Se		act ctg gct ctc Thr Leu Ala Leu 620	•
			ggc ggt gga aag Gly Gly Gly Lys 635	
			gtg tog got ctc Val Ser Ala Leu	
			agg agc act cac Arg Ser Thr His 670	ccc cac 2016
	Cys lle lle Pr		aaa act cct ggt Lys Thr Pro Gly 685	
gag cat gag	ctt gtc ctg ca	t cag ctg agg	tgt aac ggt gtg Cys Asn Gly Val	

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Gly	He				Arg	Lys				Ser	Arg				gca	2160	
705 gac		aaa	cag	aga	710		gtg	tta	aat	715 gca		ect	atc	cct	720 gaa	. 2208	
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ury	uiii	rne	740	кър	Ser	Lys	Lys	745	ser	GIU	Lys	Leu	750		ser		
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Phe	Lys 770	Ala	Gly	Leu	Leu	Gly 775	Leu	Leu	Glu	Glu	Met 780	Arg	Asp	Giu	Lys		
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785					790					795	-		·		800		
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Cys	He	Gin	Tyr 820	Asn	Val	Arg		Phe 825	Met	Asn	Val	Lys	His 830	Trp	Pro		
												aaa Lys				2544	

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Asn Lys Val Lys Asn Leu Thr Glu Glu Met Ala Gly Leu Asp Glu Thr

98	80	985	990	
	tg acc aag gag a eu Thr Lys Glu Ly 10		_	3024
	gat gac ctg cag	gca gag gag gag	c aaa gtc aac acc c Lys Val Asn Thr 1020	3069
		-	a gtg gat gat ctt n Val Asp Asp Leu 1035	3114
			g atg gat cta gaa g Met Asp Leu Glu ,	. 3159
-			ttg gct caa gaa Leu Ala Gin Giu 1065	3204
			ctt gat gaa aag Leu Asp Glu Lys 1080	3249
			caa agc aag att Gin Ser Lys ile 1095	3294
	aa gcc ctt ggt (In Ala Leu Gly (1105		aag aaa atc aag Lys Lys IIe Lys 1110	3339
	cc cgc att gag (la Arg lle Glu (gaa atc gag gca Glu ile Glu Ala	3384

	1115	i				1120)				1125	5				
gag	cgg	gco	tco	cgg	gcc	aaa	gca	gag	aag	cag	cgc	tct	gat	ctc		3429
Glu	Arg	Ala	Ser	Arg	Ala	Lys	Ala	Glu	Lys	Gin	Arg	Ser	Asp	Leu		
	1130)				1135	i				1140)				
tcc	cgg	gag	ctg	gag	gag	atc	agt	gag	agg	ctg	gaa	gaa	gcc	ggt		3474
Ser	Arg	Glu	Leu	Glu	Glu	He	Ser	Glu	Arg	Leu	Glu	Glu	Ala	Gly		
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ggg	gçc	acc	tcg	gcc	cag	att	gag	atg	aac	aag	aag	cgg	gaa	gct	:	3519
Gly	Ala	Thr	Ser	Ala	Gln	He	Glu	Met	Asn	Lys	Lys	Arg	Glu	Ala		
	1160					1165					1170					
gag	ttc	cag	aaa	atg	cgc	agg	gac	ctg	gag	gag	gcc	acc	cta	cag		3564
Glu	Phe	Gln	Lys	Met	Arg	Arg	Asp	Leu	Glu	Glu	Ala	Thr	Leu	Gin		•
	1175					1180					1185					
cat	gag	gcc	acg	gcg	gcc	acc	ctg	agg	aag	aag	cat	gca	gat	agt		3609
His	Glu	Ala	Thr	Ala	Ala	Thr	Leu	Arg	Lys	Lys	His	Ala	Asp	Ser		
	1190					1195					1200					
gtg	gcc	gag	ctt	ggg	gag	cag	att	gac	aac	ctg	cag	cga	gtg	aag		3654
Vai	Ala	Glu	Leu	Gly	Glu	Gln	He	Asp	Asn	Leu	Gln	Arg	Val	Lys		
	1205					1210					1215					
cag	aag	ctg	gag	aag	gag	aag	agt	gag	atg	aag	atg	gag	atc	gat		3699
Gln	Lys	Leu	Glu	Lys	Glu	Lys	Ser	Glu	Met	Lys	Met	Glu	He	Asp		
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Asp	Leu	Ala	Ser	Asn	Met	Glu	Thr	Val	Ser	Lys	Ala	Lys	Gly	Asn		
	1235					1240					1245		-			
															٠	
ctt	gaa	aag	atg	tgc	cgc	gct	cta	gaa	gat	caa	ctg	agt	gaa	att		3789
Leu						Ala						Ser				

	1250					1255					1260)				
aag	acc	aag	gaa	gag	gag	cag	cag	cgg	ctg	atc	aat	gac	ctc	aca		3834
Lys	Thr	Lys	Glu	Glu	Ģlu	Gln	GIn	Arg	Leu	He	Asn	Asp	Leu	Thr		
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gca	cag	aga	gcg	cgc	ctg	caa	aca	gaa	tca	ggt	gaa	tat	tca	cgc		3879
Ala	Gin	Arg	Ala	Arg	Leu	Gln	Thr	Glu	Ser	Gly	Glu	Tyr	Ser	Arg		
	1280					1285					1290				:	
	cta														:	3924
GIn	Leu	Asp	Glu	Lys	Asp		Leu	Val	Ser	Gin		Ser	Arg	Giy		
	1295					1300					1305		•			
	caa															3969
Lys	Gln	Ala	rne	ınr	GIN		He	Glu	Glu	Leu	-	Arg	Gin	Leu	١,	-)
	1310					1315					1320					
500	~~~	~~~	a+a													404.4
	gag												_	_		4014
uiu	Glu 1325	aiu	116	Lys	АТА	Lys	Ser	АІА	Leu	Ala		ма	Leu	GIN		
	1323					1330					1335					
tee																
	tee	cac	cat	тас	+~+	gae.	o+ a	ot a	000	700	005	+-+	~~~	~~~		40E0
Ser	tcc Ser													-		4059
Ser	Ser					Asp					Gin			-		4059
Ser														-		4059
	Ser 1340	Arg	His	Asp	Cys	Asp 1345	Leu	Leu	Arg	Glu	GIn 1350	Tyr	Glu	Glu		
gag	Ser 1340 cag	Arg gaa	His	Asp aag	Cys	Asp 1345 gag	Leu cta	Leu cag	Arg aga	Glu gca	Gin 1350 atg	Tyr	Glu aag	Glu	·	4059 4104
gag	Ser 1340 cag GIn	Arg gaa	His	Asp aag	Cys	Asp 1345 gag Glu	Leu cta	Leu cag	Arg aga	Glu gca	Gin 1350 atg Met	Tyr	Glu aag	Glu		
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gag Glu aac	Ser 1340 cag GIn	Arg gaa Glu gag	gcc Ala gtt	Asp aag Lys gcc	gcc Ala	Asp 1345 gag Glu 1360 tgg	Cta Ceu	Cag Gin	Arg aga Arg aaa	gca Ala	GIn 1350 atg Met 1365 gag	Tyr tcc Ser	Glu aag Lys gat	gcc Ala		
gag Glu aac	Ser 1340 cag Gln 1355 agt	Arg gaa Glu gag	gcc Ala gtt	Asp aag Lys gcc	gcc Ala	Asp 1345 gag Glu 1360 tgg	Cta Ceu	Cag Gin	Arg aga Arg aaa	gca Ala	GIn 1350 atg Met 1365 gag	tcc Ser aca Thr	Glu aag Lys gat	gcc Ala		4104
gag Glu aac	Ser 1340 cag Gln 1355 agt Ser	Arg gaa Glu gag	gcc Ala gtt	Asp aag Lys gcc	gcc Ala	Asp 1345 gag Glu 1360 tgg Trp	Cta Ceu	Cag Gin	Arg aga Arg aaa	gca Ala	GIn 1350 atg Met 1365 gag Glu	tcc Ser aca Thr	Glu aag Lys gat	gcc Ala		4104
gag Glu aac Asn	Ser 1340 cag Gln 1355 agt Ser	gaa Glu gag Glu	gcc Ala gtt Val	aag Lys gcc Ala	gcc Ala cag Gin	Asp 1345 gag Glu 1360 tgg Trp 1375	cta Leu agg Arg	cag Gin acc Thr	aga Arg aaa Lys	gca Ala tat Tyr	GIn 1350 atg Met 1365 gag Glu 1380	tcc Ser aca Thr	Glu aag Lys gat Asp	gcc Ala gcc Ala		4104

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cag	cgt	ctg	cag	gat	gct	gag	gaa	cat	gta	gaa	gct	gte	; aat	gcc	4239
Gln	Arg	Leu	Gln	Asp	Ala	Glu	Glu	His	Val	Glu	Ala	Val	Asn	Ala	
	1400	1				1405					1410)			
aaa	tgt	gct	tcc	ctt	gag	aag	acg	aag	cag	agg	ctc	cag	aat	gaa	4284
Lys	Cys	Ala	Ser	Leu	Glu	Lys	Thr	Lys	Gin	Arg	Leu	Gln	Asn	Glu	
	1415				•	1420					1425				•
gtt	gag	gac	ctc	atg	att	gat	gtt	gag	agg	aca	aat	gct	gcc	tgt	4329
Val	Glu	Asp	Leu	Met	He	Asp	Val	Glu	Arg	Thr	Asn	Ala	Ala	Cys	
•	1430					1435					1440			-	
gcc	gcc	ctg	gac	aaa	aag	caa	agg	aac	ttt	gat	aag	atc	ctg	gca	4374
Ala	Ala	Leu	Asp	Lys	Lys	Gln	Arg	Asn	Phe	Asp	Lys	He	Leu	Ala	1, 0
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gaa	tgg	aaa	cag	aag	tgt	gaa	gaa	act	cat	gct	gaa	ctt	gaa	gct	4419
Glu	Trp	Lys	Gin	Lys	Cys	Glu	Glu	Thr	His	Ala	Glu	Leu	Glu	Ala	
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Ser	Gln	Lys	Glu	Ser	Arg	Ser	Leu	Ser	Thr	Glu	Leu	Phe	Lys	He	
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Lys	Asn	Ala	Tyr	Glu	Glu	Ser	Leu	Asp	Gin	Leu	Glu	Thr	Leu	Lys	
	1490					1495					1500				
cgg	gaa	aat	aag	aat	ctg	cag	cag	gag	att	tct	gat	ctc	act	gaa	4554
Arg	Glu	Asn	Lys	Asn	Leu	Gin	Gln	Glu	He	Ser	Asp	Leu	Thr	Glu	
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Gin	He	Ala	Glu	Gly	Gly	Lys	Arg	He	His	Glu	Leu	Glu	Lys	lle	

	1520)		152	5			1530	0			
		Glr		a gaa n Glu 1540	Lys				Ala			4644
		Ala		ctt Leu 1555	Glu				Lys	_		4689
	atc lle 1565	GIn		aac Asn 1570	Gln				Val		:	4734
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_				tcc Ser 1600								4824
	agg Arg 1610			gcc Ala 1615						_		4869
				gaa Glu 1630								4914
Met				agg Arg 1645			Asn	acc				4959
ctc	aag			cac His			gct	ctc	cgg Arg			5004

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	g ctg u Leu 168!	Glr			Glu		act Thr 1695	Leu		_		5094
	g gag r Glu 1700	Arg			Ala		ctc Leu 1710	Leu		-	÷	5139
_	t gaa r Glu 1715	Arg			His		acc Thr 1725				r,	.5184
•	e acc n Thr 1730	Lys			Thr		caa Gln 1740					5229
	g atg J Met 1745	Glu										5274
	; aag : Lys 1760	Lys										5319
	aag Lys 1775	Glu						atg Met				5364
	ttg Leu								_			5409 ·

	1790)				1795	;				1800)				
	gag															5454
Ala	Glu		ı Leu	Ala	Leu			Gly	Lys	Lys			Gir	ı Lys		
	1805)				1810)				1815)			•	
cte	gag	g C C	ឧទ្ទ	øtt	cet	gaa	ctt		aat	g a a	gtt	maa	ant			5499
_	Glu															5433
	1820				0	1825	•	u.u	u.,	u.u	1830		001	aiu		
cag	aag	cgc	aat	gtt	gaa	gct	., gtc	aag	ggt	cta	CgC	aaa	cat	gag		5544
	Lys														•	
	1835					1840					1845					
aga	aaa	gtg	aag	gaa	ctc	act	tac	caa	act	gag	gaa	gac	cgc	aag		5589
Arg	Lys	Val	Lys	Glu	Leu	Thr	Tyr	Gln	Thr	Glu	Glu	Asp	Arg	Lys	1.	•)
	1850					1855					1860				·	
aat	att	ctc	agg	ctg	cag	gac	ctg	gtg	gac	aag	ctg	caa	gca	aag		5634
Asn	lle	Leu	Arg	Leu	Gin	Asp	Leu	Val	Asp	Lys	Leu	Gin	Ala	Lys		
	1865					1870					1875					
	aaa															5679
Val	Lys	Ser	Tyr	Lys	Arg		Ala	Glu	Glu	Ala		Glu	Gln	Ser		
	1880					1885					1890					
											cac			_		5724
ASII	Va I 1895	ASII	Leu	ser	Lys		Arg	Arg	11e	Gin		Glu	Leu	Glu		
	1095					1900					1905					
gag	gcc	gag	gaa	agg	gct	gac	att	get	σaσ	tee	caa	ate	220	225		5760
	Ala															5769
-	1910			0		1915		4		-01	1920	+ u i	7.011	-,,0		
ctg	agg	gtg	aag	agc	agg	gag	gtt	cac	aca	aaa	atc	ata	agt	gaa		5814
	Arg													_		

1925 1930 1935

gag taa tttatctaac tgctgaaagg tgaccaaaga aatgcacaaa atgtgaaaat 5870 Glu

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Asp Ala Lys Thr Ser Val Phe Val Val Asp Pro Lys Glu Ser Phe Val 35 40 45

Lys Ala Thr Val Gin Ser Arg Giu Giy Giy Lys Val Thr Ala Lys Thr 50 55 60

Glu Ala Gly Ala Thr Val Thr Val Lys Asp Asp Gin Val Phe Pro Met 65 70 75 80 Asn Pro Pro Lys Tyr Asp Lys Ile Glu Asp Met Ala Met Met Thr His

85 90 95

Leu His Glu Pro Ala Val Leu Tyr Asn Leu Lys Glu Arg Tyr Ala Ala 100 105 110

Trp Met Ile Tyr Thr Tyr Ser Gly Leu Phe Cys Val Thr Val Asn Pro 115 120 125

Tyr Lys Trp Leu Pro Val Tyr Asn Ala Glu Val Val Thr Ala Tyr Arg 130 135 140

Gly Lys Lys Arg Gln Glu Ala Pro Pro His IIe Phe Ser IIe Ser Asp 145 150 155 160

Asn Ala Tyr Gin Phe Met Leu Thr Asp Arg Glu Asn Gin Ser ile Leu 165 170 175

lle Thr Gly Glu Ser Gly Ala Gly Lys Thr Val Asn Thr Lys Arg Val 180 185 190

lle Gin Tyr Phe Ala Thr ile Ala Vai Thr Gly Glu Lys Lys Lys Glu 195 200 205

Glu Val Thr Ser Gly Lys Met Gln Gly Thr Leu Glu Asp Gln lle lle 210 215 220 Ser Ala Asn Pro Leu Leu Glu Ala Phe Gly Asn Ala Lys Thr Val Arg 225 230 235 240

Asn Asp Asn Ser Ser Arg Phe Gly Lys Phe IIe Arg IIe His Phe Gly
245 250 255

Thr Thr Gly Lys Leu Ala Ser Ala Asp IIe Glu Thr Tyr Leu Leu Glu 260 265 270

Lys Ser Arg Val Thr Phe Gin Leu Lys Ala Glu Arg Ser Tyr His lie 275 280 285

Phe Tyr Gln lle Met Ser Asn Lys Lys Pro Asp Leu lle Glu Met Leu 290 295 300

Leu lle Thr Thr Asn Pro Tyr Asp Tyr Ala Phe Val Ser Gln Gly Glu 305 310 315 320

lie Thr Val Pro Ser lie Asp Asp Gin Glu Glu Leu Met Ala Thr Asp 325 330 335

Ser Ala IIe Glu IIe Leu Gly Phe Thr Ser Asp Glu Arg Val Ser IIe 340 345 350

Tyr Lys Leu Thr Gly Ala Val Met His Tyr Gly Asn Met Lys Phe Lys 355 360 365 Gin Lys Gin Arg Giu Giu Gin Ala Giu Pro Asp Giy Thr Giu Vai Ala 370 375 380

Asp Lys Ala Ala Tyr Leu Gin Asn Leu Asn Ser Ala Asp Leu Leu Lys 385 390 395 400

Ala Leu Cys Tyr Pro Arg Val Lys Val Gly Asn Glu Tyr Val Thr Lys
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Gly Gln Thr Val Gln Gln Val Tyr Asn Ala Val Gly Ala Leu Ala Lys 420 425 430

Ala Val Tyr Asp Lys Met Phe Leu Trp Met Val Thr Arg IIe Asn Gin 435 440 445

Gin Leu Asp Thr Lys Gin Pro Arg Gin Tyr Phe IIe Gly Val Leu Asp 450 455 460

Ile Ala Gly Phe Glu Ile Phe Asp Phe Asn Ser Leu Glu Gln Leu Cys 465 470 475 480

Ile Asn Phe Thr Asn Glu Lys Leu Gln Gln Phe Phe Asn His His Met
485 490 495

Phe Val Leu Glu Glu Glu Glu Tyr Lys Lys Glu Gly lle Glu Trp Thr
500 505 510

Phe lle Asp Phe Gly Met Asp Leu Ala Ala Cys lle Glu Leu lle Glu
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Lys Pro Met Gly lle Phe Ser lle Leu Glu Glu Glu Cys Met Phe Pro . 530 535 540

Lys Ala Thr Asp Thr Ser Phe Lys Asn Lys Leu Tyr Glu Gln His Leu 545 550 555 560

Gly Lys Ser Asn Asn Phe Gin Lys Pro Lys Pro Ala Lys Gly Lys Pro 565 570 575

Glu Ala His Phe Ser Leu IIe His Tyr Ala Gly Thr Val Asp Tyr Asn 580 585 590

Ile Ala Gly Trp Leu Asp Lys Asn Lys Asp Pro Leu Asn Glu Thr Val
595 600 605

Val Gly Leu Tyr Gln Lys Ser Ala Met Lys Thr Leu Ala Leu Leu Phe 610 615 620

Val Gly Ala Thr Gly Ala Glu Ala Glu Ala Gly Gly Gly Lys Lys Gly 625 630 635 640

Gly Lys Lys Gly Ser Ser Phe Gln Thr Val Ser Ala Leu Phe Arg 645 650 655 Glu Asn Leu Asn Lys Leu Met Thr Asn Leu Arg Ser Thr His Pro His 660 665 670

Phe Val Arg Cys IIe IIe Pro Asn Glu Thr Lys Thr Pro Gly Ala Met . 675 680 685

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Asp Phe Lys Gln Arg Tyr Lys Val Leu Asn Ala Ser Ala lle Pro Glu
725 730 735

Gly Gln Phe IIe Asp Ser Lys Lys Ala Ser Glu Lys Leu Gly Ser 740 745 750

Ile Asp Ile Asp His Thr Gln Tyr Lys Phe Gly His Thr Lys Val Phe
755 760 765

Phe Lys Ala Gly Leu Leu Gly Leu Leu Glu Glu Met Arg Asp Glu Lys
770 780

Leu Ala Gin Leu IIe Thr Arg Thr Gin Ala Met Cys Arg Giy Phe Leu 785 790 795 800

Ala Arg Val Glu Tyr Gln Lys Met Val Glu Arg Arg Glu Ser Ile Phe 805 810 815

Cys lle Gln Tyr Asn Val Arg Ala Phe Met Asn Val Lys His Trp Pro . 820 825 830

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Met Val Thr Leu Met Gin Glu Lys Asn Asp Leu Gin Leu Gin Val Gin 885 890 895

Ala Glu Ala Asp Ser Leu Ala Asp Ala Glu Glu Arg Cys Asp Gln Leu 900 905 910

lie Lys Thr Lys lie Gin Leu Glu Aia Lys lie Lys Glu Val Thr Glu 915 920 925

Arg Ala Glu Asp Glu Glu IIe Asn Ala Glu Leu Thr Ala Lys Lys 930 935 940 Arg Lys Leu Glu Asp Glu Cys Ser Glu Leu Lys Lys Asp IIe Asp Asp 945 950 955 960

Leu Glu Leu Thr Leu Ala Lys Val Glu Lys Glu Lys His Ala Thr Glu .
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Asn Lys Val Lys Asn Leu Thr Glu Glu Met Ala Gly Leu Asp Glu Thr 980 985 990

11e Ala Lys Leu Thr Lys Glu Lys Lys Ala Leu Gln Glu Ala His Gln 995 1000 1005

Gin Thr Leu Asp Asp Leu Gin Ala Giu Giu Asp Lys Vai Asn Thr 1010 1015 1020

Leu Thr Lys Ala Lys !le Lys Leu Glu Gln Gln Val Asp Asp Leu 1025 1030 1035

Glu Gly Ser Leu Glu Gln Glu Lys Lys lie Arg Met Asp Leu Glu 1040 1045 1050

Arg Ala Lys Arg Lys Leu Glu Gly Asp Leu Lys Leu Ala Gln Glu 1055 1060 1065

Ser Ala Met Asp IIe Glu Asn Asp Lys Gln Gln Leu Asp Glu Lys 1070 1075 1080 Leu Lys Lys Glu Phe Glu Met Ser Gly Leu Gln Ser Lys lie 1085 1090 1095

Glu Asp Glu Gin Ala Leu Gly Met Gin Leu Gin Lys Lys Ile Lys
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Glu Leu Gln Ala Arg lle Glu Glu Leu Glu Glu Glu lle Glu Ala 1115 1120 1125

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Ser Arg Glu Leu Glu Glu IIe Ser Glu Arg Leu Glu Glu Ala Gly 1145 1150 1155 1. 9

Gly Ala Thr Ser Ala Gln IIe Glu Met Asn Lys Lys Arg Glu Ala 1160 1165 1170

Glu Phe Gln Lys Met Arg Arg Asp Leu Glu Glu Ala Thr Leu Gln 1175 1180 1185

His Glu Ala Thr Ala Ala Thr Leu Arg Lys Lys His Ala Asp Ser 1190 1195 1200

Val Ala Glu Leu Gly Glu Gin lle Asp Asn Leu Gin Arg Val Lys 1205 1210 1215 Gin Lys Leu Giu Lys Giu Lys Ser Giu Met Lys Met Giu lie Asp 1220 1225 1230

Asp Leu Ala Ser Asn Met Glu Thr Val Ser Lys Ala Lys Gly Asn 1235 1240 1245

Leu Glu Lys Met Cys Arg Ala Leu Glu Asp Gln Leu Ser Glu IIe 1250 1255 1260

Lys Thr Lys Glu Glu Glu Gln Gln Arg Leu lle Asn Asp Leu Thr 1265 1270 1275

Ala Gin Arg Ala Arg Leu Gin Thr Giu Ser Giy Giu Tyr Ser Arg 1280 1285 1290

Gin Leu Asp Giu Lys Asp Thr Leu Vai Ser Gin Leu Ser Arg Giy 1295 1300 1305

Lys Gin Ala Phe Thr Gin Gin lie Giu Giu Leu Lys Arg Gin Leu 1310 1315 1320

Glu Glu Ile Lys Ala Lys Ser Ala Leu Ala His Ala Leu Gln 1325 1330 1335

Ser Ser Arg His Asp Cys Asp Leu Leu Arg Glu Gln Tyr Glu Glu 1340 1345 1350

- Glu Gln Glu Ala Lys Ala Glu Leu Gln Arg Ala Met Ser Lys Ala 1355 1360 1365
- Asn Ser Glu Val Ala Gln Trp Arg Thr Lys Tyr Glu Thr Asp Ala 1370 1375 1380
- lle Gin Arg Thr Giu Giu Leu Giu Giu Ala Lys Lys Lys Leu Ala 1385 1390 1395
- Gln Arg Leu Gln Asp Ala Glu Glu His Val Glu Ala Val Asn Ala 1400 . 1405 1410
- Lys Cys Ala Ser Leu Glu Lys Thr Lys Gln Arg Leu Gln Asn Glu 1415 1420 1425
- Val Glu Asp Leu Met ile Asp Val Glu Arg Thr Asn Ala Ala Cys 1430 1435 1440
- Ala Ala Leu Asp Lys Lys Gln Arg Asn Phe Asp Lys IIe Leu Ala 1445 1450 1455
- Glu Trp Lys Gln Lys Cys Glu Glu Thr His Ala Glu Leu Glu Ala 1460 1465 1470
- Ser Gin Lys Giu Ser Arg Ser Leu Ser Thr Giu Leu Phe Lys IIe 1475 1480 1485

Lys Asn Ala Tyr Glu Glu Ser Leu Asp Gln Leu Glu Thr Leu Lys 1490 1495 1500

Arg Glu Asn Lys Asn Leu Gln Gln Glu IIe Ser Asp Leu Thr Glu
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Gin He Ala Giu Giy Giy Lys Arg He His Giu Leu Giu Lys He 1520 1525 1530

Lys Lys Gin Val Glu Gin Glu Lys Ser Glu Leu Gin Ala Ala Leu 1535 1540 1545

Glu Glu Ala Glu Ala Ser Leu Glu His Glu Glu Gly Lys lle Leu 1550 1555 1560

Arg lie Gin Leu Giu Leu Asn Gin Vai Lys Ser Giu Vai Asp Arg 1565 1570 1575

Lys lle Ala Glu Lys Asp Glu Glu lle Asp Gln Met Lys Arg Asn 1580 1585 1590

His lie Arg lie Val Glu Ser Met Gin Ser Thr Leu Asp Ala Glu 1595 1600 1605

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Leu Lys Asp Thr Gln Leu His Leu Asp Asp Ala Leu Arg Ser Gln 1655 1660 1665

Glu Asp Leu Lys Glu Gln Leu Ala Met Val Glu Arg Arg Ala. Asn 1670 1675 1680

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Ser Glu Arg Val Gln Leu Leu His Thr Gln Asn Thr Ser Leu lle 1715 1720 1725

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Glu Met Glu Asp IIe IIe Gln Glu Ala Arg Asn Ala Glu Glu Lys 1745 1750 1755 Ala Lys Lys Ala Ile Thr Asp Ala Ala Met Met Ala Glu Glu Leu 1760 1765 1770

Lys Lys Glu Gln Asp Thr Ser Ala His Leu Glu Arg Met Lys Lys
1775 1780 1785

Asn Leu Glu Gln Thr Val Lys Asp Leu Gln His Arg Leu Asp Glu 1790 1795 1800

Ala Glu Gln Leu Ala Leu Lys Gly Gly Lys Lys Gln IIe Gln. Lys 1805 1810 1815

Leu Glu Ala Arg Val Arg Glu Leu Glu Gly Glu Val Glu Ser Glu 1820 1825 1830

Gin Lys Arg Asn Val Glu Ala Val Lys Gly Leu Arg Lys His Glu
1835 1840 1845

Arg Lys Val Lys Glu Leu Thr Tyr Gln Thr Glu Glu Asp Arg Lys 1850 1855 1860

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Val Lys Ser Tyr Lys Arg Gin Ala Giu Giu Ala Giu Giu Gin Ser 1880 1885 1890 Asn Val Asn Leu Ser Lys Phe Arg Arg Ile Gln His Glu Leu Glu 1895 1900 1905

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Met Leu Gin Thr Lys Asp

1 5

ctc atc tgg act ttg ttt ttc ctg gga act gca gtt tct ctg cag gtg

Leu lle Trp Thr Leu Phe Phe Leu Gly Thr Ala Val Ser Leu Gln Val

10 15 20

gat att gtt ccc agc cag ggg gag atc agc gtt gga gag tcc aaa ttc 151

Asp	lle	Va I 25	Pro	Ser	Gln	Gly	Glu 30	ile	Ser	Val	Gly	G I u 35	Ser	Lys	Phe		
ttc	tta	tgc	caa	gtg	gca	gga	gat	gcc	aaa	gat	aaa	gac	atc	tcc	tgg		199
Phe	Leu 40	Cys	Gin	Val	Ala	Gly 45	Asp	Ala	Lys	Asp	Lys 50	Asp	lle	Ser	Trp	•	
ttc	tcc	CCC	aat	gga	gaa	aag	ctc	acc	cca	aac	cag	cag	cgg	atc	tca		247
Phe	Ser	Pro	Asn	Gly	Glu	Lys	Leu	Thr	Pro	Asn	Gin	Gin	Arg	lle	Ser	•	
55					60					65					70		
gtg	gtg	tgg	aat	gat	gat	tcc	tcc	tcc	acc	ctc	acc	atc	tat	aac	gcc	:	295
Val	Val	Trp	Asn	Asp	Asp	Ser	Ser	Ser	Thr	Leu	Thr	He	Tyr	Asn	Ala		
				75					80					85			
aac	atc	gac	gac	gcc	ggc	att	tac	aag	tgt	gtg	gtt	aca	ggc	gag	gat	,	343
·Asn	He	Asp	Asp	Ala	Gly	He	Tyr	Lys	Cys	Val	Val	Thr	Gly	Glu	Asp		
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ggc	agt	gag	tca	gag	gcc	acc	gtc	aac	gtg	aag	atc	ttt	cag	aag	ctc		391
Gly	Ser		Ser	Glu	Ala	Thr	Val	Asn	Val	Lys	He	Phe	Gin	Lys	Leu		
		105					110					115					
atg	ttc	aag	aat	gcg	cca	acc	cca	cag	gag	ttc	cgg	gag	ggg	gaa	gat		439
Met		Lys	Asn	Ala	Pro	Thr	Pro	Gln	Glu	Phe	Arg	Glu	Gly	Glu	Asp		
	120					125					130						
gcc	gtg	att	gtg	tgt	gat	gtg	gtc	agc	tcc	ctc	cca	cca	acc	atc	atc		487
Ala	Val	He	Val	Cys	Asp	Val	Val	Ser	Ser	Leu	Pro	Pro	Thr	He	He		
135					140					145					150		
			aaa														535
Trp	Lys	His	Lys		Arg	Asp	Val	He		Lys	Lys	Asp	Val		Phe		
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ata	gtc	ctg	tcc	aac	aac	tac	ctg	cag	atc	cgg	ggc	atc	aag	aaa	aca		583

	ile	val	Leu	Ser 170		Asn	Tyr	Leu	G1n 175		Arg	; Gly	lle	Lys 180	_	Thr		
				Thr					Gly					Arg		gag Glu		631
-			ttc Phe										Pro					679
	•	Ala	agg Arg												_		•	727
			ctg Leu													agc Ser	•. *	775
			aag Lys															823
			ttc Phe 265												-	-		871
			gac Asp						_		Ala			_	_			919
			gat Asp		Thr					Val		_						967
	aca	tat	gta	gag	aac	cag	act	gcc	atg	gaa	tta	gag	gag	cag	gtc	act		1015

WO 2005/012512 90/206 PCT/JP2004/011401

Thr	· Tyı	- Va	l Glu	315		n Thr	- Ala	a Met	320		Glı	u Glu	ı Gir	n Va∣ 325	Thr		
ctt	aco	; tg	t gaa	gco	tco	gga	gad	ccc	att	ccc	to	c ato	acc	; tgg	agg		1063
Leu	Thr	· Cy:	330		ı Ser	Gly	Asp	335		Pro	Sei	r lle	340	·	Arg	•	
												ctg					1111
Thr	Ser			Asn	He	Ser			Glu	Lys	Thr	Leu		Gly	His	•	
		345	•				350)				355					
atg	gtg	gte	cgt	agc	cat	gcc	cgt	gtg	tcg	tcg	ctg	acc	ctg	aag	agc		1159
Met			Arg	Ser	His	Ala	Arg	Val	Ser	Ser	Leu	Thr	Leu	Lys	Ser		
	360					365					370)					
atc	cag	tac	act	gat	gcc	gga	gag	tac	atc	tgc	acc	gcc	agc	aac	acc	, ,	1207
												Ala				••	
375					380					385					390		
	~~~						•										
												caa Gin					1255
	,	٠	пор	395	<b>u</b>	00,	mo L	• yı	400	uiu	Vai	uiii	ıyr	405	Pro		
												ggg				1	303
Lys	Leu	GIn		Pro	Val	Ala	Val		Thr	Trp	Glu	Gly	Asn	Gln	Val		
			410					415					420				
aac	atc	acc	tgc	gag	gta	ttt	gcc	tat	ccc	agt	gcc	acg	atc	tca	tgg	1	351
Asn	He	Thr	Cys	Glu	Val	Phe	Ala	Tyr	Pro	Ser	Ala	Thr	Пе	Ser	Trp		
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ttt	cgg	gat	ggc	cag	ctg	ctg	cca	agc	tcc	aat	tac	agc	aat	atc		1	399
												Ser			_	•	555
	440					445					450				-		
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lle 455		Asn	Thr	Pro	Ser 460		Ser	Tyr	Leu	G1u 465		Thr	Pro	Asp	Ser 470		
			•		Asn					Ala					ggg Gly	<b>1495</b>	
												acc Thr				1543 ·	
												gcc Ala 515			_	1591	
												ctc Leu			aaa Lys	, √1639	
												tcc Ser	_			1687	
			Glu									atc He	_		_	1735	
												ctc Leu				1783	
						Ala					Lys	acg Thr 595	_		_	1831	
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Gln	Gly 600		Pro	Ser	Ala	Pro 605		Leu	Giu	Gly	GIn 610		Gly	Glu	Asp		
gga	aac	tct	att	aaa	gtg	aac	ctg	atc	aag	cag	gat	gac	ggo	ggc	tcc		1927
Gly 615		Ser	lle	Lys	Va I 620	Asn	Leu	lle	Lys	G1n 625	•	Asp	Gly	Gly	Ser 630	•	
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Pro	116	Arg	ніѕ	635	Leu	va i	Arg	lyr	Arg 640	Ala	Leu	Ser	Ser	645	Trp		
		gag Glu													aag Lvs	:	2023
			650	_				655					660		_,_		
		gac Asp														١.	<b>∘2071</b>
		665					670					675					
		gga Gly												_	_	٠	2119
	680					685					690						
		aca Thr															2167
695	.,,	••••		110	700	AIG	Aon	uly	061	705	7111	961	uly	Leu	710		
		gcc										_	_		_		2215
Thr	Gly	Ala	lle	715	Gly	He	Leu	lle	Va I 720	lle	Phe	Val	Leu	Leu 725	Leu		
		gtg												_			2263
Val	Vai	Val	Asp 730	He	Thr	Cys	Tyr	Phe 735	Leu	Asn	Lys	Cys	Gly 740	Leu	Phe		
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Met	Cys	11e 745	Ala	Val	Asn	Leu	Cys 750	Gly	Lys	Ala	Gly	Pro 755	Gly	Ala	Lys	-
	Lys					Gly	_	_	gcc Ala		_		_			<b>2359</b>
	760					765					770			÷		
aag	gag	ccc	atc	gtg	gag	gtt	cga	acg	gag	gag	gag	agg	acc	cca	aac	2407
	Glu	Pro	He	Val		Val	Arg	Thr	Glu		Glu	Arg	Thr	Pro		•
775					780					785					790	
cat	gat	gga	ggg	aaa	cac	aca	gag	ccc	aac	gag	acc	acg	cca	ctg	acg	· 2455
His	Asp	Gly	Gly		His	Thr	Glu	Pro		Glu	Thr	Thr	Pro	Leu	Thr	
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gag	ccc	gag	aag	ggc	ccc	gta	gaa	gca	aag	cca	gag	tgc	cag	gag	aca	. →2503
Glu	Pro	Glu	Lys	Gly	Pro	Val	Glu	Ala	Lys	Pro	Glu	Cys	Gln	Glu	Thr	
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gaa	acg	aag	cca	gcg	cca	gcc	gaa	gtc	aag	acg	gtc	CCC	aat	gac	gcc	2551
Glu	Thr	Lys	Pro	Ala	Pro	Ala	Glu	Val	Lys	Thr	Val	Pro	Asn	Asp	Ala	
		825					830				·	835				
aca	cag	aca	aag	gag	aac	gag	agc	aaa	gca	tga	tggg	tgaa	ga g	aaco	gagca	2604
Thr	Gln	Thr	Lys	Glu	Asn	Glu	Ser	Lys	Ala							
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<210> 8

**<211> 848** 

<212> PRT

<213≻ Homo sapiens

<400> 8

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Ala Val Ser Leu Gin Val Asp lie Val Pro Ser Gin Gly Glu lie Ser 20 25 30

Val Gly Glu Ser Lys Phe Phe Leu Cys Gln Val Ala Gly Asp Ala Lys 35 40 45

Asp Lys Asp IIe Ser Trp Phe Ser Pro Asn Gly Glu Lys Leu Thr Pro 50 55 60

Asn Gin Gin Arg ile Ser Val Val Trp Asn Asp Asp Ser Ser Ser Thr
65 70 75 80

Leu Thr lie Tyr Asn Ala Asn lie Asp Asp Ala Gly lie Tyr Lys Cys
85 90 95

Val Val Thr Gly Glu Asp Gly Ser Glu Ser Glu Ala Thr Val Asn Val
100 105 110

Lys lie Phe Gin Lys Leu Met Phe Lys Asn Ala Pro Thr Pro Gin Giu 115 120 125

Phe Arg Glu Gly Glu Asp Ala Val IIe Val Cys Asp Val Val Ser Ser 130 135 140 Leu Pro Pro Thr lle lle Trp Lys His Lys Gly Arg Asp Val lle Leu 145 150 155 160

Lys Lys Asp Val Arg Phe IIe Val Leu Ser Asn Asn Tyr Leu Gln IIe 165 170 175

Arg Gly lie Lys Lys Thr Asp Glu Gly Thr Tyr Arg Cys Glu Gly Arg 180 185 190

lie Leu Ala Arg Gly Glu lie Asn Phe Lys Asp ile Gin Val lie Val 195 200 205

Asn Val Pro Pro Thr IIe Arg Ala Arg Gin Asn IIe Val Asn Ala Thr 210 215 220

Ala Asn Leu Gly Gln Ser Val Thr Leu Val Cys Asp Ala Glu Arg Phe 225 230 235 240

Pro Glu Pro Thr Met Ser Trp Thr Lys Asp Gly Glu Gln ile Glu Gln
245 250 255

Glu Glu Asp Asp Glu Lys Tyr lle Phe Ser Asp Asp Ser Ser Gln Leu 260 265 270

Thr lie Lys Lys Val Asp Lys Asn Asp Glu Ala Glu Tyr lie Cys lie 275 280 285 Ala Glu Asn Lys Ala Gly Glu Gln Asp Ala Thr lie His Leu Lys Val 290 295 300

Phe Ala Lys Pro Lys ile Thr Tyr Vai Glu Asn Gln Thr Ala Met Glu 305 310 315 320

Leu Glu Glu Gln Val Thr Leu Thr Cys Glu Ala Ser Gly Asp Pro IIe 325 330 335

Pro Ser IIe Thr Trp Arg Thr Ser Thr Arg Asn IIe Ser Ser Glu Glu
340 345 350

Lys Thr Leu Asp Gly His Met Val Val Arg Ser His Ala Arg Val Ser 355 360 365

Ser Leu Thr Leu Lys Ser IIe Gin Tyr Thr Asp Ala Giy Giu Tyr IIe 370 375 380

Cys Thr Ala Ser Asn Thr lie Gly Gln Asp Ser Gln Ser Met Tyr Leu 385 390 395 400

Glu Val Gln Tyr Ala Pro Lys Leu Gln Gly Pro Val Ala Val Tyr Thr 405 410 415

Trp Glu Gly Asn Gln Val Asn He Thr Cys Glu Val Phe Ala Tyr Pro
420 425 430

Ser Ala Thr lie Ser Trp Phe Arg Asp Gly Gln Leu Leu Pro Ser Ser
435
440
445

Asn Tyr Ser Asn IIe Lys IIe Tyr Asn Thr Pro Ser Aia Ser Tyr Leu 450 455 460

Glu Val Thr Pro Asp Ser Glu Asn Asp Phe Gly Asn Tyr Asn Cys Thr 465 470 475 480

Ala Val Asn Arg lie Gly Gin Glu Ser Phe Glu Phe lie Leu Val Gin 485 490 495

Ala Asp Thr Pro Ser Ser Pro Ser IIe Asp Gln Val Glu Pro Tyr Ser 500 505 510

Ser Thr Ala Gin Val Gin Phe Asp Glu Pro Glu Ala Thr Gly Gly Val 515 520 525

Pro Ile Leu Lys Tyr Lys Ala Glu Trp Arg Ala Val Gly Glu Glu Val 530 535 540

Trp His Ser Lys Trp Tyr Asp Ala Lys Glu Ala Ser Met Glu Gly 11e 545 550 555 560

Val Thr IIe Val Gly Leu Lys Pro Glu Thr Thr Tyr Ala Val Arg Leu 565 570 575

Ala Ala Leu Asn Gly Lys Gly Leu Gly Glu IIe Ser Ala Ala Ser Glu 580 585 590

Phe Lys Thr Gln Pro Val Gln Gly Glu Pro Ser Ala Pro Lys Leu Glu 595 600 605

Gly Gln Met Gly Glu Asp Gly Asn Ser lie Lys Val Asn Leu lie Lys 610 615 . 620

Gin Asp Asp Gly Gly Ser Pro !le Arg His Tyr Leu Val Arg Tyr Arg 625 630 635 640

Ala Leu Ser Ser Glu Trp Lys Pro Glu IIe Arg Leu Pro Ser Gly Ser 645 650 655

Asp His Val Met Leu Lys Ser Leu Asp Trp Asn Ala Glu Tyr Glu Val 660 665 670

Tyr Val Val Ala Glu Asn Gln Gln Gly Lys Ser Lys Ala Ala His Phe 675 680 685

Val Phe Arg Thr Ser Ala Gin Pro Thr Ala IIe Pro Ala Asn Giy Ser 690 695 700

Pro Thr Ser Gly Leu Ser Thr Gly Ala IIe Val Gly IIe Leu IIe Val
705 710 715 720

Ile Phe Val Leu Leu Leu Val Val Val Asp ile Thr Cys Tyr Phe Leu
725 730 735

Asn Lys Cys Gly Leu Phe Met Cys IIe Ala Val Asn Leu Cys Gly Lys
740 745 750

Ala Gly Pro Gly Ala Lys Gly Lys Asp Met Glu Glu Gly Lys Ala Ala 755 760 765

Phe Ser Lys Asp Glu Ser Lys Glu Pro IIe Val Glu Val Arg Thr Glu 770 780

Glu Glu Arg Thr Pro Asn His Asp Gly Gly Lys His Thr Glu Pro Asn 785 790 795 800

Glu Thr Thr Pro Leu Thr Glu Pro Glu Lys Gly Pro Vai Glu Ala Lys 805 810 815

Pro Glu Cys Gln Glu Thr Glu Thr Lys Pro Ala Pro Ala Glu Val Lys 820 825 830

Thr Val Pro Asn Asp Ala Thr Gln Thr Lys Glu Asn Glu Ser Lys Ala 835 840 845

<210> 9

<211> 1692

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> (121).. (1080)

**<400> 9** 

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gccactctct gccgcttggg ttgggcgaaa gccaggaccg tgccgcgcca ccgccaggat 120

atg gag cta ctg tcg cca ccg ctc cgc gac gta gac ctg acg gcc ccc

Met Glu Leu Leu Ser Pro Pro Leu Arg Asp Val Asp Leu Thr Ala Pro

1 10 15

gac ggc tct ctc tgc tcc ttt gcc aca acg gac gac ttc tat gac gac

216

Asp Gly Ser Leu Cys Ser Phe Ala Thr Thr Asp Asp Phe Tyr Asp Asp

20 25 30

ccg tgt ttc gac tcc ccg gac ctg cgc ttc ttc gaa gac ctg gac ccg 264

Pro Cys Phe Asp Ser Pro Asp Leu Arg Phe Phe Glu Asp Leu Asp Pro

5 40 4

cgc ctg atg cac gtg ggc gcg ctc ctg aaa ccc gaa gag cac tcg cac

Arg Leu Met His Val Gly Ala Leu Leu Lys Pro Glu Glu His Ser His

50 55 60

ttc ccc gcg gcg gtg cac ccg gcc ccg ggc gca cgt gag gac gag cat

Phe Pro Ala Ala Val His Pro Ala Pro Gly Ala Arg Glu Asp Glu His

65 70 75 80

gtg cgc gcg ccc agc ggg cac cac cag gcg ggc cgc tgc cta ctg tgg 408

Val Arg Ala Pro Ser Gly His His Gln Ala Gly Arg Cys Leu Leu Trp 85 90 95

															aag		456
АІА	Uys	Lys		Cys	Lys	Arg	Lys			Asn	Ala	ı Asp			Lys		
			100					105					110				
<b>"</b> 00	<b>400</b>	200	ot a	0.00	g o g	0.77	000	0.50	a+~								504
															gcc	•	504
ліа	Ala	115	me L	AI g	uiu	Aig	120		Leu	ડલા	Lys		ASII	Giu	Ala		
		113					120					125					
ttt	gag	aca	ctc	aag	cgc	tgc	acg	tcg	agc	aat	cca	aac	cag	cgg	ttg		552
Phe	Glu	Thr	Leu	Lys	Arg	Cys	Thr	Ser	Ser	Asn	Pro	Asn	Gln	Arg	Leu		
	130					135					140						
															•	٠	
ccc	aag	gtg	gag	atc	ctg	cgc	aac	gcc	atc	cgc	tat	atc	gag	ggç	ctg		600
Pro	Lys	Val	Glu	He	Leu	Arg	Asn	Ala	lle	Arg	Tyr	lle	Glu	Gly	Leu		
145					150					155					160		
																٠,٠	
cag	gct	ctg	ctg	cgc	gac	cag	gac	gcc	gcg	CCC	cct	ggc	gca	gcc	gcc	•	648
Gin	Ala	Leu	Leu	Arg	Asp	Gln	Asp	Ala	Ala	Pro	Pro	Gly	Ala	Ala	Ala		
-				165					170					175			
												ggc					696
Phe	Tyr	Ala		Gly	Pro	Leu	Pro	Pro	Gly	Arg	Gly	Gly	Glu	His	Tyr		
			180					185					190				
						_											
												tgc					744
Ser	GIY		Ser	Asp	Ala	Ser		Pro	Arg	Ser	Asn	Cys	Ser	Asp	Gly		
		195					200					205					
at «	at a	<b>700</b>	+00	000	~~~	222									4		700
												cgg					792
M€ L	210	nsp	ıyı	Sei			F1 0	oer	шу			Arg	Arg	ASI	Cys		
	210					215					220						
tac	gaa	ggc	acc .	tac	tac	aac	gag	āCa	വാ	agc	раа	CCC	agg	cee.	aaa		840
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225					230		u	, 🟎		235			- " Б		240		
															~ <del>~</del> ~		

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cgc atc tcc acc gag agc cct gcg gcg ccc gcc ctc ctg ctg gcg gac  Arg lle Ser Thr Glu Ser Pro Ala Ala Pro Ala Leu Leu Leu Ala Asp  260 265 270	936
gtg cct tct gag tcg cct ccg cgc agg caa gag gct gcc gcc ccc agc Val Pro Ser Glu Ser Pro Pro Arg Arg Gln Glu Ala Ala Pro Ser 275 280 285	984
gag gga gag agc agc ggc gac ccc acc cag tca ccg gac gcc gcc ccg Glu Gly Glu Ser Ser Gly Asp Pro Thr Gln Ser Pro Asp Ala Ala Pro 290 295 300	1032
cag tgc cct gcg ggt gcg aac ccc aac ccg ata tac cag gtg ctc tga Gin Cys Pro Ala Giy Ala Asn Pro Asn Pro Ile Tyr Gin Val Leu 305 310 315	1080
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gctaatttat aa 1692

<210> 10

<211> 319

<212> PRT

<213> Homo sapiens

<400> 10

Met Glu Leu Leu Ser Pro Pro Leu Arg Asp Val Asp Leu Thr Ala Pro

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Asp Gly Ser Leu Cys Ser Phe Ala Thr Thr Asp Asp Phe Tyr Asp Asp 20 25 30

Pro Cys Phe Asp Ser Pro Asp Leu Arg Phe Phe Glu Asp Leu Asp Pro 35 40 45

Arg Leu Met His Val Gly Ala Leu Leu Lys Pro Glu Glu His Ser His 50 55 60

Phe Pro Ala Ala Val His Pro Ala Pro Gly Ala Arg Glu Asp Glu His 65 70 75 80

Val Arg Ala Pro Ser Gly His His Gln Ala Gly Arg Cys Leu Leu Trp 85 90 95 Ala Cys Lys Ala Cys Lys Arg Lys Thr Thr Asn Ala Asp Arg Lys
100 105 110

Ala Ala Thr Met Arg Glu Arg Arg Leu Ser Lys Val Asn Glu Ala 115 120 125

Phe Glu Thr Leu Lys Arg Cys Thr Ser Ser Asn Pro Asn Gln Arg Leu
130 135 140

Pro Lys Val Glu IIe Leu Arg Asn Ala IIe Arg Tyr IIe Glu Gly Leu

145 150 155 160

Gln Ala Leu Leu Arg Asp Gln Asp Ala Ala Pro Pro Gly Ala Ala Ala 165 170 175

Phe Tyr Ala Pro Gly Pro Leu Pro Pro Gly Arg Gly Gly Glu His Tyr 180 185 190

Ser Gly Asp Ser Asp Ala Ser Ser Pro Arg Ser Asp Cys Ser Asp Gly 195 200 205

Met Met Asp Tyr Ser Gly Pro Pro Ser Gly Ala Arg Arg Arg Asn Cys 210 215 220

Tyr Glu Gly Ala Tyr Tyr Asn Glu Ala Pro Ser Glu Pro Arg Pro Gly
225 230 235 240

Lys Ser Ala Ala Val Ser Ser Leu Asp Tyr Leu Ser Ser ile Val Glu 245 250 255

Arg Ile Ser Thr Glu Ser Pro Ala Ala Pro Ala Leu Leu Leu Ala Asp 260 265 270

Val Pro Ser Glu Ser Pro Pro Arg Arg Gln Glu Ala Ala Ala Pro Ser 275 280 285

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Gin Cys Pro Ala Gly Ala Asn Pro Asn Pro Ile Tyr Gin Val Leu 305 310 315

⟨210⟩ 11

<211> 1427

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

⟨222⟩ (43).. (810)

**<400> 11** 

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Met Asp Val Met

54

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Asp	Gly	Cys	Gin	Phe	Ser	Pro	Ser	Glu	Tyr	Phe	Tyr	Asp	Gly	Ser	Cys		
5					10					15					20		
ata	ccg	tcc	ccc	gag	ggt	gaa	ttt	ggg	gac	gag	ttt	gtg	ccg	cga	gtg		150
He	Pro	Ser	Pro	Glu	Gly	Glu	Phe	Gly	Asp	Glu	Phe	Val	Pro	Arg	Val		
				25					30					35			
												•				•	
			gga														198
Ala	Ala	Phe	Gly	Ala	His	Lys	Ala		Ļeu	Gln	Gly	Ser		Glu	Asp	:	
			40					45					50				
~~~																	0.40
			cga														246
GIU	піѕ	55	Arg	AIA	rro	Ittr		піѕ	піѕ	uin	Ala		піѕ	Gys	Leu	•	
		55					60					65			•	'. '	
atø	†øø	acc	tgc	222	acc	tøc	220	200	220	tcc	acc	200	at a	σat	cau		294
			Cys														234
	70		0,0	_,0	,,,,	75	_,0	/·'' 5	_,0	001	80	****	1110 C	nop	AI B		
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cgg	aag	gca	gcc	act	atg		gag	cgg	agg	cgc		aag	aag	gtc	aac		342
			gcc Ala			cgc					ctg						342
						cgc					ctg						342
Arg					Met	cgc				Arg	ctg				Asn		342
Arg 85	Lys	Ala		Thr	Met 90	cgc Arg	Glu	Arg	Arg	Arg 95	ctg Leu	Lys	Lys	Val	Asn 100		342 390
Arg 85 cag	Lys gct	Ala ttc	Ala	Thr acc	Met 90 ctc	cgc Arg aag	Glu agg	Arg tgt	Arg acc	Arg 95 acg	ctg Leu acc	Lys aac	Lys	Va I aac	Asn 100 cag		
Arg 85 cag	Lys gct	Ala ttc	Ala gaa	Thr acc	Met 90 ctc	cgc Arg aag	Glu agg	Arg tgt	Arg acc	Arg 95 acg	ctg Leu acc	Lys aac	Lys	Va I aac	Asn 100 cag		
Arg 85 cag	Lys gct	Ala ttc	Ala gaa	Thr acc Thr	Met 90 ctc	cgc Arg aag	Glu agg	Arg tgt	Arg acc Thr	Arg 95 acg	ctg Leu acc	Lys aac	Lys	Val aac Asn	Asn 100 cag		
Arg 85 cag Gin	Lys gct Ala	Ala ttc Phe	Ala gaa	Thr acc Thr 105	Met 90 ctc Leu	cgc Arg aag Lys	Glu agg Arg	Arg tgt Cys	Arg acc Thr 110	Arg 95 acg Thr	ctg Leu acc Thr	Lys aac Asn	Lys ccc Pro	Val aac Asn 115	Asn 100 cag Gin		
Arg 85 cag Gln	Lys gct Ala ctg	ttc Phe	Ala gaa Glu	Thr acc Thr 105 gtg	Met 90 ctc Leu gag	cgc Arg aag Lys	Glu agg Arg	Arg tgt Cys	acc Thr 110	Arg 95 acg Thr	ctg Leu acc Thr	Lys aac Asn	CCC Pro	Val aac Asn 115	Asn 100 cag Gin		390
Arg 85 cag Gln	Lys gct Ala ctg	ttc Phe	gaa Glu aag	Thr acc Thr 105 gtg	Met 90 ctc Leu gag	cgc Arg aag Lys	Glu agg Arg	Arg tgt Cys	acc Thr 110	Arg 95 acg Thr	ctg Leu acc Thr	Lys aac Asn	CCC Pro	Val aac Asn 115	Asn 100 cag Gin		390
Arg 85 cag Gln	Lys gct Ala ctg	ttc Phe	gaa Glu aag Lys	Thr acc Thr 105 gtg	Met 90 ctc Leu gag	cgc Arg aag Lys	Glu agg Arg	tgt Cys agg Arg	acc Thr 110	Arg 95 acg Thr	ctg Leu acc Thr	Lys aac Asn	CCC Pro tac Tyr	Val aac Asn 115	Asn 100 cag Gin		390
Arg 85 cag Gin agg Arg	gct Ala ctg Leu	ttc Phe ccc Pro	gaa Glu aag Lys	Thr acc Thr 105 gtg Val	Met 90 ctc Leu gag Glu	cgc Arg aag Lys atc	agg Arg ctc Leu	tgt Cys agg Arg 125	acc Thr 110 aat Asn	Arg 95 acg Thr gcc Ala	ctg Leu acc Thr	aac Asn cgc Arg	ccc Pro tac Tyr 130	aac Asn 115 atc	Asn 100 cag Gin gag Giu		390
Arg 85 cag Gin agg Arg	gct Ala ctg Leu	ttc Phe ccc Pro	gaa Glu aag Lys 120	Thr acc Thr 105 gtg Val	Met 90 ctc Leu gag Glu ctg	cgc Arg aag Lys atc Ile	agg Arg ctc Leu	tgt Cys agg Arg 125	acc Thr 110 aat Asn	Arg 95 acg Thr gcc Ala	ctg Leu acc Thr	aac Asn cgc Arg	ccc Pro tac Tyr 130	aac Asn 115 atc Ile	Asn 100 cag Gin gag Glu		390 438

								acc Thr						_		534
	150					155					160					
gat	ggc	atg	ccc	gaa	tgt	aac	agt	cct	gtc	tgg	tcc	aga	aag	agc	agt	582
Asp	Gly	Met	Pro	Glu	Cys	Asn	Ser	Pro	Val	Trp	Ser	Arg	Lys	Ser	Ser	
165					170					175					180	•
act	ttt	gac	agc	atc	tac	tgt	cct	gat	gta	tca	aat	gta	tat	gcc	aca	630
Thr	Phe	Asp	Ser	lle	Tyr	Cys	Pro	Asp	Val	Ser	Asn	Val	Tyr	Ala	Thr	:
				185					190					195		
gat	aaa	aac	tcc	tta	tcc	agc	ttg	gat	tgc	tta	tcc	aac	ata	gtg	gac	678
Asp	Lys	Asn	Ser	Leu	Ser	Ser	Leu	Asp	Cys	Leu	Ser	Asn	He	Val	Asp	
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cgg	atc	acc	tcc	tca	gag	caa	cct	ggg	ttg	cct	ctc	cag	gat	ctg	gct	726
Arg	lle	Thr	Ser	Ser	Glu	Gln	Pro	Gly	Leu	Pro	Leu	Gin	Asp	Leu	Ala	
-		215					220					225				
tct	ctc	tct	cca	gtt	gcc	agc	acc	gat	tca	cag	cct	cga	act	cca	ggg	774
Ser	Leu	Ser	Pro	Val	Ala	Ser	Thr	Asp	Ser	GIn	Pro	Arg	Thr	Pro	Gly	
	230					235					240					
gct	tct	agt	tcc	agg	ctt	atc	tat	cat	gtg	cta	tga	acta	attt	tc		820
Ala	Ser	Ser	Ser	Arg	Leu	He	Tyr	His	Val	Leu						
245					250					255						
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tccc	aaac	ca a	gaca	acat	g ta	cata	aaga	ttt	cttt	tca	gttg	taaa	ıtt t	gtaa	agatt	940
acct	tgcc	ac t	ttat	aaga	ıa ag	tgta	ttta	act	aaaa	agt	cato	attg	ca a	ataa	tactt	1000
tctt	cttc	tt t	atta	ttct	t tg	ctta	gata	tta	atac	ata	gtto	cagt	aa t	acta	tttct	1060

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taaaacatta	aaacagctga	gaatcagtta	aatggaattt	taaatatatt	taactatttc	1240
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tatattttga	tottttcttg	taagaaatgt	atcttttaaa	tgtaagcaca	aaatagtact	1360
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<211> 255

<212> PRT

<213> Homo sapiens

⟨400⟩ 12

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Val Pro Arg Val Ala Ala Phe Gly Ala His Lys Ala Glu Leu Gln Gly 35 40 45

Ser Asp Glu Asp Glu His Val Arg Ala Pro Thr Gly His His Gln Ala

50

55

60

Gly His Cys Leu Met Trp Ala Cys Lys Ala Cys Lys Arg Lys Ser Thr 65 70 75 80

Thr Met Asp Arg Arg Lys Ala Ala Thr Met Arg Glu Arg Arg Leu 85 90 95

Lys Lys Val Asn Gin Ala Phe Glu Thr Leu Lys Arg Cys Thr Thr Thr 100 105 110 .

Asn Pro Asn Gin Arg Leu Pro Lys Vai Giu ile Leu Arg Asn Ala ile , , , 115 120 125

Arg Tyr lle Glu Ser Leu Gin Glu Leu Leu Arg Glu Gin Val Glu Asn 130 135 140

Tyr Tyr Ser Leu Pro Gly Gln Ser Cys Ser Glu Pro Thr Ser Pro Thr 145 150 155 160

Ser Asn Cys Ser Asp Gly Met Pro Glu Cys Asn Ser Pro Val Trp Ser 165 170 175

Arg Lys Ser Ser Thr Phe Asp Ser IIe Tyr Cys Pro Asp Val Ser Asn 180 185 190

Val Tyr Ala Thr Asp Lys Asn Ser Leu Ser Ser Leu Asp Cys Leu Ser

195 200 205

Asn lie Val Asp Arg lie Thr Ser Ser Glu Gln Pro Gly Leu Pro Leu 210 215 220

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<211> 675 ·

<212> DNA

<213> Homo sapiens

<220>

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1 5 10 15

tat gat ggg gaa aac tac ctg cct gtc cac ctc cag ggc ttc gaa cca

Tyr Asp Gly Glu Asn Tyr Leu Pro Val His Leu Gln Gly Phe Glu Pro

20 25 30

cca ggc tac gag cgg acg gag ctc acc ctg agc ccc gag gcc cca ggg
Pro Gly Tyr Glu Arg Thr Glu Leu Thr Leu Ser Pro Glu Ala Pro Gly

ccc ctt gag gac aag ggg ctg ggg acc ccc gag cac tgt cca ggc cag Pro Leu Glu Asp Lys Gly Leu Gly Thr Pro Glu His Cys Pro Gly Gin tgc ctg ccg tgg gcg tgt aag gtg tgt aag agg aag tcg gtg tcc gtg Cys Leu Pro Trp Ala Cys Lys Val Cys Lys Arg Lys Ser Val Ser Val gac cgg cgg cgg gcc aca ctg agg gag aag cgc agg ctc aag aag Asp Arg Arg Arg Ala Ala Thr Leu Arg Glu Lys Arg Arg Leu Lys Lys 95. gtg aat gag gcc ttc gag gcc ctg aag aga agc acc ctg ctc aac ccc Val Asn Glu Ala Phe Glu Ala Leu Lys Arg Ser Thr Leu Leu Asn Pro aac cag cgg ctg ccc aag gtg gag atc ctg cgc agt gcc atc cag tac Asn Gin Arg Leu Pro Lys Val Giu ile Leu Arg Ser Ala ile Gin Tyr atc gag cgc ctc cag gcc ctg ctc agc tcc ctc aac cag gag gag cgt lle Glu Arg Leu Gln Ala Leu Leu Ser Ser Leu Asn Gln Glu Glu Arg gac ctc cgc tac cgg ggc ggg ggc ggg ccc cag cca ggg gtg ccc agc Asp Leu Arg Tyr Arg Gly Gly Gly Gly Pro Gln Pro Gly Val Pro Ser gaa tgc agc tct cac agc gcc tcc tgc agt cca gag tgg ggc agt gca Glu Cys Ser Ser His Ser Ala Ser Cys Ser Pro Glu Trp Gly Ser Ala ctg gag ttc agc gcc aac cca ggg gat cat ctg ctc acg gct gac cct

Leu Glu Phe Ser Ala Asn Pro Gly Asp His Leu Leu Thr Ala Asp Pro

180 185 190

aca gat gcc cac aac ctg cac tcc ctc acc tcc atc gtg gac agc atc

624

Thr Asp Ala His Asn Leu His Ser Leu Thr Ser Ile Val Asp Ser Ile

195

200

205

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Thr Val Glu Asp Val Ser Val Ala Phe Pro Asp Glu Thr Met Pro Asp

210

220

tag . 675

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<212> PRT

<213> Homo sapiens

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Pro Gly Tyr Glu Arg Thr Glu Leu Thr Leu Ser Pro Glu Ala Pro Gly
35 40 45

Pro Leu Glu Asp Lys Gly Leu Gly Thr Pro Glu His Cys Pro Gly Gln
50 55 60

Cys Leu Pro Trp Ala Cys Lys Val Cys Lys Arg Lys Ser Val Ser Val 65 70 75 80

Asp Arg Arg Ala Ala Thr Leu Arg Glu Lys Arg Arg Leu Lys Lys
85 90 95

Val Asn Glu Ala Phe Glu Ala Leu Lys Arg Ser Thr Leu Leu Asn Pro
100 105 110

Asn Gln Arg Leu Pro Lys Val Glu IIe Leu Arg Ser Ala IIe Gln Tyr 115 120 125

lle Glu Arg Leu Gin Ala Leu Leu Ser Ser Leu Asn Gin Glu Glu Arg 130 135 140

Asp Leu Arg Tyr Arg Gly Gly Gly Pro Gln Pro Gly Val Pro Ser 145 150 155 160

Glu Cys Ser Ser His Ser Ala Ser Cys Ser Pro Glu Trp Gly Ser Ala 165 170 175

Leu Glu Phe Ser Ala Asn Pro Gly Asp His Leu Leu Thr Ala Asp Pro 180 185 190

Thr Asp Ala His Asn Leu His Ser Leu Thr Ser IIe Val Asp Ser IIe 195 200 205 Thr Val Glu Asp Val Ser Val Ala Phe Pro Asp Glu Thr Met Pro Asn 210 215 220

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<213> Homo sapiens

⟨220⟩

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<222> (373).. (1902)

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cag gag aag ggc ctg tcc ggc gcc ccc agc ccc acc atg tcc gag gac

459
Gin Glu Lys Gly Leu Ser Gly Ala Pro Ser Pro Thr Met Ser Glu Asp

15
20
25

tcc gcg ggc tcg ccc tgc ccg tcg ggc tcc ggc tcg gac acc gag aac 507

	Se 30		la	Gly	/ Se	r Pr	o Cy 35	rs Pr	o Se	er Gi	y Se	r GI 40		er As	sp Th	ır G	lu As 45		.•
							u As	c ac			o Ly	s GI				p Le	eu Ly		555
						g ga	g ga	c aag				g tgo					g gt		603
					65				•	70		٠.		٠.	· 75				• .
			n V					tac Tyr											651
								aag Lys 100	Asn					Lys					· , → 699
								gcg)	747
								aac Asn									Lys		795
				g l				gag Glu											843
				g L				cag Gin											891
1	tac	cag	CC	g c	gg (cgg	agg	aag ·	tcg	gte	aag	aac :	o o o	cag	aca	asa	uca.		020

Tyı	Gln	Pro	Arg	Arg	Arg	Lys	Ser	Val	Lys	Asn	Gly	Gin	Ala	Glu	Ala	
	175					180					185					
													_			
															aag	987
	ı Glu `	Ala	ınr	uiu			ніѕ	116	Ser			Ala	116	Phe	_	•
190	,				195					200					205	
gcg	g ctg	cag	gcc	gac	tcg	cca	cac	tcc	tcc	tcc	ggc	atg	agc	gag	gtg	1035
	Leu								•							·
				210				•	215		•			220		
																:
	tcc															1083
His	Ser	Pro		Glu	His	Ser	Gly	Gln	Ser	GIn	Gly	Pro	Pro	Thr	Pro	
			225					230					235			
000		222	000													
	Thr														aag	,, → 1131
110	' '''	240		Lys	1111	weh	245	uiii	FIU	uly	LyS	250	ASP	Leu	Lys	
		240					270					250				
cga	gag	ggg	cgc	CCC	ttg	cca	gag	ggg	ggc	aga	cag	CCC	cct	atc	gac	1179
Arg	Glu	Gly	Arg	Pro	Leu	Pro	Glu	Gly	Gly	Arg	Gin	Pro	Pro	lle	Asp	
	255					260					265					
	cgc															1227
	Arg	Asp	Val	Asp		Gly	Glu	Leu	Ser	Ser	Asp	Val	He	Ser		
270					275					280					285	
ato	gag	200	++0	got	gt o	000	~~~	+++	~~~		+	a+~				1075
	Glu															1275
	u,u		1110	290	141	AOII	aru	1 110	295	uiii	ı yı	LGu	110	300	ASII	
ggc	cac	ccg	ggg	gtg	ccg	gcc	acg	cac	ggc	cag	gtc	acc	tac	acg	ggc	1323
	His															
			305					310					315			
agc	tac	ggc	atc	agc	agc	acc	gcg	gcc	acc	ccg	gcg	agc	gcg	ggc	cac	1371

Ser	Tyr	Gly 320	lle	Ser	Ser	Thr	Ala 325	Ala	Thr	Pro	Ala	Ser 330	Ala	Gly	His	
			tcc Ser												_	1419
			ccg Pro			_										1467
			cag GIn													1515
_			ctg Leu 385	_	_		_			_						→1563
			cag GIn													1611
			cag GIn				_								_	1659
			ccg Pro				_				_			_		1707
			agc Ser													1755
ctc	tac	tcc	acc	ttc	acc	tac	atg	aac	ccc	gct	cag	cgc	CCC	atg	tac	1803

WO 2005/012512 118/206 PCT/JP2004/011401

Leu Tyr Ser Thr Phe Thr Tyr Met Asn Pro Ala Gin Arg Pro Met Tyr 465 470 475 acc ccc atc gcc gac acc tct ggg gtc cct tcc atc ccg cag acc cac 1851 Thr Pro Ile Ala Asp Thr Ser Gly Val Pro Ser Ile Pro Gln Thr His 480 485 490 agc ccc cag cac tgg gaa caa ccc gtc tac aca cag ctc act cga cct 1899 Ser Pro Gln His Trp Glu Gln Pro Val Tyr Thr Gln Leu Thr Arg Pro 495 500 505 tga ggaggcctcc cacgaagggc gaagatggcc gagatgatcc taaaaataac 1952 cgaagaaaga gaggaccaac cagaattccc tttggacatt tgtgtttttt tgtttttta 2012 ttttgttttg tttttcttc ttcttcttct tccttaaaga catttaagct aaaggcaact, 32072 cgtacccaaa tttccaagac acaaacatga cctatccaag cgcattaccc acttgtggcc 2132 aatcagtggc caggccaacc ttggctaaat ggagcagcga aatcaacgag aaactggact 2192 ttttaaaccc tcttcagagc aagcgtggag gatgatggag aatcgtgtga tcagtgtgct 2252 aaatctctct gcctgtttgg actttgtaat tatttttta gcagtaatta aagaaaaaag 2312 toctctgtga ggaatattct ctattttaaa tatttttagt atgtactgtg tatgattcat 2372 taccattttg aggggattta tacatatttt tagataaaat taaatgctct tatttttcca 2432 acagctaaac tactcttagt tgaacagtgt gccctagctt ttcttgcaac cagagtattt 2492 2552 aaaacagaat tgtgttatgt gatcagtttt ggggggttaac tttgcttaat tcctcaggct 2612 ttgcgattta aggaggagct gccttaaaaa aaaataaagg ccttattttg caattatggg 2672

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aaa 3935

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<211> 509

<212> PRT

<213> Homo sapiens

<400> 16

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Gly Leu Ser Gly Ala Pro Ser Pro Thr Met Ser Glu Asp Ser Ala Gly
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Ser Pro Cys Pro Ser Gly Ser Gly Ser Asp Thr Glu Asn Thr Arg Pro 35 40 45

Gin Glu Asn Thr Phe Pro Lys Gly Glu Pro Asp Leu Lys Lys Glu Ser 50 55 60

Glu Glu Asp Lys Phe Pro Val Cys IIe Arg Glu Ala Val Ser Gln Val 65 70 75 80 Leu Lys Gly Tyr Asp Trp Thr Leu Val Pro Met Pro Val Arg Val Asn 85 90 95

Gly Ser Ser Lys Asn Lys Pro His Val Lys Arg Pro Met Asn Ala Phe 100 105 110

Met Val Trp Ala Gin Ala Ala Arg Arg Lys Leu Ala Asp Gin Tyr Pro 115 120 125

His Leu His Asn Ala Glu Leu Ser Lys Thr Leu Gly Lys Leu Trp Arg 130 135 140

1. 4

Leu Leu Asn Glu Ser Glu Lys Arg Pro Phe Val Glu Glu Ala Glu Arg 145 150 155 160

Leu Arg Val Gin His Lys Lys Asp His Pro Asp Tyr Lys Tyr Gin Pro 165 170 175

Arg Arg Arg Lys Ser Val Lys Asn Gly Gin Ala Glu Ala Glu Glu Ala 180 185 190

Thr Glu Gln Thr His IIe Ser Pro Asn Ala IIe Phe Lys Ala Leu Gln 195 200 205

Ala Asp Ser Pro His Ser Ser Ser Gly Met Ser Glu Val His Ser Pro 210 215 220 Gly Glu His Ser Gly Gln Ser Gln Gly Pro Pro Thr Pro Pro Thr Thr
225 230 235 240

Pro Lys Thr Asp Val Gin Pro Gly Lys Ala Asp Leu Lys Arg Glu Gly
245 250 255

Arg Pro Leu Pro Glu Gly Gly Arg Gln Pro Pro IIe Asp Phe Arg Asp 260 265 270

Val Asp IIe Gly Glu Leu Ser Ser Asp Val IIe Ser Asn IIe Glu Thr 275 280 285

Phe Asp Val Asn Glu Phe Asp Gln Tyr Leu Pro Pro Asn Gly His Pro 290 295 300

Gly Val Pro Ala Thr His Gly Gln Val Thr Tyr Thr Gly Ser Tyr Gly 305 310 315 320

lle Ser Ser Thr Ala Ala Thr Pro Ala Ser Ala Gly His Val Trp Met 325 330 335

Ser Lys Gln Gln Ala Pro Pro Pro Pro Gln Gln Pro Pro Gln Ala 340 345 350

Pro Pro Ala Pro Gin Ala Pro Pro Gin Pro Gin Ala Ala Pro Pro Gin 355 360 365 Gin Pro Ala Ala Pro Pro Gin Gin Pro Gin Ala His Thr Leu Thr Thr 370 375 380

Leu Ser Ser Glu Pro Gly Gln Ser Gln Arg Thr His IIe Lys Thr Glu 385 390 395 400

Gin Leu Ser Pro Ser His Tyr Ser Giu Gin Gin Gin His Ser Pro Gin
405 410 415

Gin Ile Ala Tyr Ser Pro Phe Asn Leu Pro His Tyr Ser Pro Ser Tyr 420 425 430

Pro Pro Ile Thr Arg Ser Gln Tyr Asp Tyr Thr Asp His Gln Asn Ser 435 440 445

Ser Ser Tyr Tyr Ser His Ala Ala Gly Gln Gly Thr Gly Leu Tyr Ser 450 455 460

Thr Phe Thr Tyr Met Asn Pro Ala Gin Arg Pro Met Tyr Thr Pro IIe 465 470 475 480

Ala Asp Thr Ser Gly Val Pro Ser IIe Pro Gln Thr His Ser Pro Gln
485 490 495

His Trp Glu Gin Pro Val Tyr Thr Gln Leu Thr Arg Pro 500 505

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tgo	atga	ggg	cgcg	gtag	ag a	cccg	gaco	c gc	gccg	tgct	cct	gccg	ttt	cgct	gcgctc	120
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cgc	ccgg	gcc	cggc	tcag	cc a	ggcc	ccgc	g gt	gago	c at	g at	t cg	c ct	c gg	g gct	175
										Me	t II	e Ar	g Le	u GI	y Ala	
_										1				5		
CCC	cag	tcg	ctg	gtg	ctg	ctg	acg	ctg	ctc	gtc	gcc	gct	gtc	ctt	cgg	223
Pro	Gin	Ser	Leu	Val	Leu	Leu	Thr	Leu	Leu	Val	Ala	Ala	Val	Leu	Arg	
			10					15					20			
tgt	cag	ggc	cag	gat	gtc	cag	gag	gct	ggc	agc	tgt	gtg	cag	gat	ggg	271
Cys	Gin	Gly	Gin	Asp	Val	Gin	Glu	Ala	Gly	Ser	Cys	Val	Gin	Asp	Gly	
		25					30					35				
cag	agg	tat	aat	gat	aag	gat	gtg	tgg	aag	ccg	gag	ccc	tgc	cgg	atc	319
Gin	Arg	Tyr	Asn	Asp	Lys	Asp	Val	Trp	Lys	Pro	Glu	Pro	Cys	Arg	He	
	40					45					50					
tgt	gtc	tgt	gac	act	ggg	act	gtc	ctc	tgc	gac	gac	ata	atc	tgt	gaa	367
Cys	Vai	Cys	Asp	Thr	Gly	Thr	Val	Leu	Cys	Asp	Asp	ile	He	Cys	Glu	
55					60					65					70	

gac	gt	g aa	a ga	ac tg	c ct	c ag	c cc	t ga	g at	c cc	c tt	c gg	a ga	ag t	tgc	tgc		415
Asp	Va	l Ly	's As	эр Су	s Le	u Se	r Pr	o GI	u II	e Pr	o Ph	e GI	y G	lu (Cys	Cys		
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					ccc Pro			•	2335
					ggc Gly				2383
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					gtt Val 770				2479
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ggc	cgc	gtt	gga	ccc	cca	ggc	tcc	aat	ggc	aac	cct	gga	ccc	cct	ggt	2911
Gly	Arg	Val	Gly	Pro	Pro	Gly	Ser	Asn	Gly	Asn	Pro	Gly	Pro	Pro	Gly	
		905					910					915				
ccc	cct	ggt	cct	tct	gga	aaa	gat	ggt	CCC	aaa	ggt	gct	cga	gga	gac	2959
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					Gln					g Gl	gc atc y lle	:	3103
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	Pro 1000	Ser			Lys				gga Gly O				3196
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	cct Pro 1030	Ala			Arg				ggt Gly)			;	3286
		Pro			Ala				ggt Gly		_	;	3331
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Gly Ser	ggc	tcc	cct	ggc	ccc	gct	ggt	cca	act	ggc	aag	caa	gga	gac	aga	3421
gga gaa gct ggt gca caa ggo ccc atg gga ccc toa gga cca gct 3466 Gly Glu Ala Gly Ala Gln Gly Pro Met Gly Pro Ser Gly Pro Ala 1100 gga gcc cgg gga atc cag ggt cct caa ggo ccc aga ggt gac aaa 3511 Gly Ala Arg Gly Ile Gln Gly Pro Gln Gly Pro Arg Gly Asp Lys 11105 gga gag gct gga gag cct ggo gag aga ggc ctg aag gga cac cgt 3556 Gly Glu Ala Gly Glu Pro Gly Glu Arg Gly Leu Lys Gly His Arg 1130 ggo tto act ggt ctg cag ggt ctg ccc ggc cct cct ggt cct tct 3601 Gly Phe Thr Gly Leu Gln Gly Leu Pro Gly Pro Pro Gly Pro Ser 3646 Gly Asp Glin Gly Ala Ser Gly Pro Ala Gly Pro Ser Gly Pro Arg 3646 Gly Pro Pro Gly Pro Val Gly Pro Ser Gly Lys Asp Gly Ala Asn 3691 gga atc cct ggc ccc att ggg cct ct ggc aaa gat ggt gct aat 3691 ggg atc cct ggc ccc att ggg cct ct ggt ccc ct ggt gga cga tca 3736	Gly	Ser	Pro	Gly	Pro	Ala	Gly	Pro	Thr	Gly	Lys	Gln	Gly	/ Asp	Arg	
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gct	ggc	tta	ggc	CCg	aga	gag	aag	ggc	CCC	gac	ccc	ctg	cag	tac	. 3871
Ala	Gly	Leu	Gly	Pro	Arg	Glu	Lys	Gly	Pro	Asp	Pro	Leu	Gin	Tyr	
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atg	cgg	gcc	gac	cag	gca	gcc	ggt	ggc	ctg	aga	cag	cat	gac	gcc	3916
Met	Arg	Ala	Asp	Gln	Ala	Ala	Gly	Gly	Leu	Arg	GIn	His	Asp	Ala	
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Glu	Val	Asp	Ala	Thr	Leu	Lys	Ser	Leu	Asn	Asn	Gin	ile	Glu	Ser	
	1255					1260					1265				
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Lys	Lys	His	s 11e	Tr) Phe	Gly	Glu	Thr	- 116	Asr	n Gly	Gly	y Phe	e His		
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Phe	Ser	Tyr	Gly	/ Asp	Asp	Asn	Leu	Ala	Pro	Asr	Thr	Ala	a Asr	n Val		
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Gin	Met	Thr	Phe	Leu	Arg	Leu	Leu	Ser	Thr	Glu	Gly	Ser	Glr	Asn		-
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	•														•	
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He	Thr	Tyr	His	Cys	Lys	Asn	Ser	He	Ala	Tyr	Leu	Asp	Glu	Ala		
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Ala	Gly	Asn	Leu	Lys	Lys	Ala	Leu	Leu	He	Gin	Gly	Ser	Asn	Asp		
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Val	He	Glu	Tyr	Arg	Ser	Gin	Lys	Thr	Ser	Arg	Leu	Pro	He	He		
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and aggree ground good too tought of the octood of the state of the st	4341
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Pro Phe Gly Glu Cys Cys Pro Ile Cys Pro Thr Asp Leu Ala Thr Ala 85 90 95

Ser Gly Gln Pro Gly Pro Lys Gly Gln Lys Gly Glu Pro Gly Asp Ile 100 105 110

Lys Asp IIe Val Gly Pro Lys Gly Pro Pro Gly Pro Gln Gly Pro Ala 115 120 125

Gly Glu Gln Gly Pro Arg Gly Asp Arg Gly Asp Lys Gly Glu Lys Gly 130 140

Ala Pro Gly Pro Arg Gly Arg Asp Gly Glu Pro Gly Thr Pro Gly Asn 145 150 155 160

Pro Gly Pro Pro Gly Pro Pro Gly Pro Pro Gly Leu Gly
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Gly Asn Phe Ala Ala Gln Met Ala Gly Gly Phe Asp Glu Lys Ala Gly 180 185 190 Gly Ala Gln Leu Gly Val Met Gln Gly Pro Met Gly Pro 195 200 205

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His Arg Gly Tyr Pro Gly Leu Asp Gly Ala Lys Gly Glu Ala Gly Ala 290 295 300

Pro Gly Val Lys Gly Glu Ser Gly Ser Pro Gly Glu Asn Gly Ser Pro 305 310 315 320

Gly Pro Met Gly Pro Arg Gly Leu Pro Gly Glu Arg Gly Arg Thr Gly 325 330 335 Pro Ala Gly Ala Ala Gly Ala Arg Gly Asn Asp Gly Gln Pro Gly Pro 340 345 350

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1. *

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Gin Gly Ala Arg Gly Gin Pro Gly Val Met Gly Phe Pro Gly Pro Lys 595 600 605

Gly Ala Asn Gly Glu Pro Gly Lys Ala Gly Glu Lys Gly Leu Pro Gly 610 615 620 Ala Pro Gly Leu Arg Gly Leu Pro Gly Lys Asp Gly Glu Thr Gly Ala 625 630 635 640

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Pro Pro Gly Glu Gly Gly Lys Pro Gly Asp Gln Gly Val Pro Gly Glu 675 680 685

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Thr Gly Pro Lys Gly Ala Arg Gly Ala Gln Gly Pro Pro Gly Ala Thr 885 890 895

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Lys Gly Ala Arg Gly Asp Ser Gly Pro Pro Gly Arg Ala Gly Glu Pro 930 935 940

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Pro Ser Gly Pro Ala Gly Ala Arg Gly lle Gln Gly Pro Gln Gly
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1. 9

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Leu Lys Gly His Arg Gly Phe Thr Gly Leu Gln Gly Leu Pro Gly
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Asp Met Ser Ala Phe Ala Gly Leu Gly Pro Arg Glu Lys Gly Pro 1220 1225 1230

Asp Pro Leu Gin Tyr Met Arg Ala Asp Gin Ala Ala Gly Gly Leu 1235 1240 1245

Arg Gln His Asp Ala Glu Val Asp Ala Thr Leu Lys Ser Leu Asn 1250 1255 1260

Asn Gin lie Giu Ser lie Arg Ser Pro Giu Giy Ser Arg Lys Asn 1265 1270 1275

Pro Ala Arg Thr Cys Arg Asp Leu Lys Leu Cys His Pro Glu Trp 1280 1285 1290

Lys Ser Gly Asp Tyr Trp IIe Asp Pro Asn Gln Gly Cys Thr Leu 1295 1300 1305

Asp Ala Met Lys Val Phe Cys Asn Met Glu Thr Gly Glu Thr Cys 1310 1315 1320 Val Tyr Pro Asn Pro Ala Asn Val Pro Lys Lys Asn Trp Trp Ser 1325 1330 1335

Ser Lys Ser Lys Glu Lys Lys His IIe Trp Phe Gly Glu Thr IIe 1340 1345 1350

Asn Gly Gly Phe His Phe Ser Tyr Gly Asp Asp Asn Leu Ala Pro 1355 1360 1365

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108

Met Thr Thr Leu Leu Trp Val Phe Val Thr Leu Arg Val IIe Thr Ala

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Ala Val Thr Val Glu Thr Ser Asp His Asp Asn Ser Leu Ser Val Ser
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204

11e Pro Gin Pro Ser Pro Leu Arg Val Leu Leu Gly Thr Ser Leu Thr

35

40

45

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															gcc Ala		5	540
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			Ser												gcc Ala			36
	gaa	gac	ggc	ttc	cac	cag	tgt	gac	gcc	ggc	tgg	ctg	gct	gac	cag	act	6	84

Glu	ı Asp	Gly 195		His	Gin	Cys	Asp 200		Gly	Trp	Leu	Ala 205		Gin	Thr	
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	Glu		cct Pro													780
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			tct Ser 260													. 3 876
_	_		ctg Leu										-		_	924
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			ggc Gly					Tyr					GIn			1068
tac	CCC	gac	ccc	tca	tcc	cgc	tac	gac	gcc	atc	tgc	tac	aca	ggt	gaa	1116

Tyr Pr	o Asp	9 Pro 340		Ser	Arg	Tyr	Asp 345		He	Cys	Tyr	Thr 350		' Glu	
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gac at Asp II 37	e Thr														1212
cct cg Pro Ar 385															1260
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Gln					Trp					Asp		tgc Cys				1932
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	_	_														
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Thr		Thr	Ser	Pro	Ser	Gly	Vai	Glu	Glu	Trp		Val	Thr	Gin	Val	
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Leu	rro		пе	ren	rro	Thr		rro	rro	inr	uly		ulu	ınr	GIU	
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	-0 -			-o-			-~u		ouu	9-8			900	Jua	946	4714

WO 2005/012512 152/206 PCT/JP2004/011401

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Ser	Glu	Glu	Pro	Phe	Pro	Ser	Val	Arg	Pro	Phe	Pro	Ser	Val	Glu	Leu	•
				805			•		8 10					815		
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Phe	Pro	Ser	Glu	Glu	Pro	Phe	Pro	Ser	Lys	Glu	Pro	Ser	Pro	Ser	Glu	
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	• • •	835		.			840	.,.	••••		00.	845		u.u		
		000					040					045				
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Asp	Leu	Asp	Ser	Ser	Gly	Leu	Thr	Ser	Thr	Val	Gly	Ser	Gly	Leu	Thr	
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gta	gaa	ag+	gge	cte	CCC	tca	aaa	σe+	gae.	asa	200	a++	ger.	+~~	000	2844
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Val Glu Ser Gly Leu Pro Ser Gly Asp Glu Glu Arg I le Glu Trp Pro 915 920 925	
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	cct Pro 1100	Gly					Ser				tct Ser 1110	Gly			: 3399
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Ala					Glu					Leu	cct Pro 1185			_	3624
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		Glu				ggg Gly 1240	Leu					Val		_	· 3804
		Ala				gag Glu 1255						Pro		_	→ 3849
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Glu	1325		· Ala	Pro	Gly	Val 1330		Glu	ılle	Ser	Gly 1335		Pro	Ser		
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Gly	Glu	Val	Leu	Glu	Thr	Ser	Thr	Ser	Ala	Val	Gly	Asp	Leu	Ser		
	1340	1				1345	i				1350	l				
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Gly			Ser	Gly	Gly			Leu	Glu	lle	Ser		Ser	Gly	•	
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Val	Glu										Val					
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Ser	Ala	Ser	Gly	He	Glu	Asp	Val	Ser	Glu	Leu	Pro	Ser	Gly	Glu		
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Gly		Glu	Ihr	Ser	Ala		Gly	Val	Glu	Asp	Leu	Ser	Arg	Leu		
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_								•			_					
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Asp	Leu	Ser	Gly	Val	Pro	Ser	Gly	Gly	Glu	Gly	Leu	Glu	Thr	Ser		
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														agg		4434
		Glu	Val	Gly			Leu	Ser	Gly		Pro	Ser	Gly	Arg		
	1445					1450					1455					
mo~	aa+	at a	~~		.	~~+	+-+					_4_				4470
Rag	88 L	uLa	gag	act	LCa	gut	LCE	gga	gCT	gag	gac	CTC	agt	ggg	4	4479

Glu	Gly 1460		Glu	Thr	Ser	Ala 1465	Ser	Gly	Ala	Glu	Asp 1470	Leu	Ser	Gly	
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	gac Asp 1490														4569
	cca Pro 1505														4614
	ggg Gly 1520											_		_	, · ·4659
	agt Ser 1535														4704
	aca Thr 1550										gca Ala 1560				4749
	ggg Gly 1565											_			4794
	ctg Leu 1580														4839
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gt o	1745		227	+++	tot	1750		et o	rot	t 00	1755				5270
_	agt										agt				5 379
Vai		ury	um	FIIE	Ser		1111	Vai	ASP	Ser	Ser	шіу	riie	III	
	1760					1765					1770				
	cag										ggc				5424
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Val	Ser	Gly	Glu	Ser	Ser	Arg	Ala	Glu	lle	Gly	Ser	Ser	Leu	Pro	
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		Thr					Gly				gta Val 1905	Ala			5784
		Gly									act Thr 1920	Leu			5829
_											att He 1935				,, →5874
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Ser											ggg Gly 1995			_	6054
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	tct Ser 2030					agc Ser 2035	Gly					Phe				6189
	gag Glu 2045					cca Pro 2050						tcc Ser			. 6	234
	agt Ser 2060					gag Glu 2065									→ 6	279
	acg Thr 2075														6	324
						ggc Gly 2095									6	369
	tct Ser 2105				Gln					Pro					6	414
	gaa Glu 2120				Ser					Ser					64	459
cac _.	aca	gtc	gaa	aca	gcc	acc	tcc	cca	aca	gat	gct	tcc	atc	cca	6	504

His	Thr 2135		Glu	Thr	Ala	Thr 2140		Pro	Thr	Asp	Ala 2145		lle	Pro	
	tct Ser 2150	Pro					Glu					Ala	_	_	6549
	gag Glu 2165	Val					Trp					Gly		_	6594
	cgc Arg 2180										gat Asp 2190	Ala			6639
	tgt Cys 2195											-			, → 6684
	gag Glu 2210										gac Asp 2220				6729
	ggc Gly 2225														6774
	gga Gly 2240				Gln										6819
Asp	aac Asn 2255				Ala					Val		atg Met			6864
cac	gag	aag	ggc	gag	tgg	aat	gat	gtt	CCC	tgc	aat	tac	cac	ctc	6909

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lle Pro Cys Tyr Phe lle Asp Pro Met His Pro Val Thr Thr Ala Pro 50 55 60

Ser Thr Ala Pro Leu Ala Pro Arg IIe Lys Trp Ser Arg Val Ser Lys
65 70 75 80

Glu Lys Glu Val Val Leu Leu Val Ala Thr Glu Gly Arg Val Arg Val 85 90 95.

Asn Ser Ala Tyr Gin Asp Lys Val Ser Leu Pro Asn Tyr Pro Ala IIe 👝 100 105 110

Pro Ser Asp Ala Thr Leu Glu Val Gln Ser Leu Arg Ser Asn Asp Ser 115 120 125

Gly Val Tyr Arg Cys Glu Val Met His Gly IIe Glu Asp Ser Glu Ala 130 135 140

Thr Leu Glu Val Val Lys Gly He Val Phe His Tyr Arg Ala He
145 150 155 160

Ser Thr Arg Tyr Thr Leu Asp Phe Asp Arg Ala Gln Arg Ala Cys Leu 165 170 175

Gin Asn Ser Ala Ile Ile Ala Thr Pro Giu Gin Leu Gin Ala Ala Tyr

180

185

190

Glu Asp Gly Phe His Gln Cys Asp Ala Gly Trp Leu Ala Asp Gln Thr 195 200 205

Val Arg Tyr Pro lle His Thr Pro Arg Glu Gly Cys Tyr Gly Asp Lys 210 215 220

Asp Glu Phe Pro Gly Val Arg Thr Tyr Gly !le Arg Asp Thr Asn Glu 225 230 235 . 240

Thr Tyr Asp Val Tyr Cys Phe Ala Giu Giu Met Giu Giy Giu Val Phe 245 250 255

Tyr Ala Thr Ser Pro Glu Lys Phe Thr Phe Gln Glu Ala Ala Asn Glu 260 265 270

Cys Arg Arg Leu Gly Ala Arg Leu Ala Thr Thr Gly His Val Tyr Leu 275 280 285

Ala Trp Gln Ala Gly Met Asp Met Cys Ser Ala Gly Trp Leu Ala Asp 290 295 300

Arg Ser Val Arg Tyr Pro IIe Ser Lys Ala Arg Pro Asn Cys Gly Gly 305 310 315 320

Asn Leu Leu Gly Val Arg Thr Val Tyr Val His Ala Asn Gln Thr Gly

325 330 335

Tyr Pro Asp Pro Ser Ser Arg Tyr Asp Ala IIe Cys Tyr Thr Gly Glu 340 345 350

Asp Phe Val Asp IIe Pro Giu Asn Phe Phe Gly Val Gly Gly Glu Glu 355 360 365

Asp lie Thr Val Gin Thr Val Thr Trp Pro Asp Met Giu Leu Pro Leu 370 380 .

Pro Arg Asn IIe Thr Glu Gly Glu Ala Arg Gly Ser Val IIe Leu Thr 385 390 395 400

Val Lys Pro IIe Phe Glu Val Ser Pro Ser Pro Leu Glu Pro Glu Glu
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Pro Phe Thr Phe Ala Pro Giu IIe Giy Ala Thr Ala Phe Ala Giu Val 420 425 430

Glu Asn Glu Thr Gly Glu Ala Thr Arg Pro Trp Gly Phe Pro Thr Pro 435 440 445

Gly Leu Gly Pro Ala Thr Ala Phe Thr Ser Glu Asp Leu Val Val Gln 450 455 460

Val Thr Ala Val Pro Gly Gln Pro His Leu Pro Gly Gly Val Val Phe

465 470 475 480

His Tyr Arg Pro Gly Pro Thr Arg Tyr Ser Leu Thr Phe Glu Glu Ala 485 490 . 495

Gin Gin Ala Cys Pro Gly Thr Gly Ala Val IIe Ala Ser Pro Glu Gin 500 505 510

Leu Gin Ala Ala Tyr Giu Ala Giy Tyr Giu Gin Cys Asp Ala Giy Trp
515 520 525

Leu Arg Asp Gin Thr Val Arg Tyr Pro IIe Val Ser Pro Arg Thr Pro 1. 530 535 540

Cys Val Gly Asp Lys Asp Ser Ser Pro Gly Val Arg Thr Tyr Gly Val 545 550 555 560

Arg Pro Ser Thr Glu Thr Tyr Asp Val Tyr Cys Phe Val Asp Arg Leu 565 570 575

Glu Gly Glu Val Phe Phe Ala Thr Arg Leu Glu Gln Phe Thr Phe Gln 580 585 590

Glu Ala Leu Glu Phe Cys Glu Ser His Asn Ala Thr Ala Thr Thr Gly 595 600 605

Gin Leu Tyr Ala Ala Trp Ser Arg Gly Leu Asp Lys Cys Tyr Ala Gly

610 615 620

Trp Leu Ala Asp Gly Ser Leu Arg Tyr Pro I le Val Thr Pro Arg Pro 625 630 635 640

Ala Cys Gly Gly Asp Lys Pro Gly Val Arg Thr Val Tyr Leu Tyr Pro 645 650 655

Asn Gln Thr Gly Leu Pro Asp Pro Leu Ser Arg His His Ala Phe Cys 660 665 670

Phe Arg Gly IIe Ser Ala Val Pro Ser Pro Gly Glu Glu Glu Gly Gly . . . 675 680 685

Thr Pro Thr Ser Pro Ser Gly Val Glu Glu Trp Ile Val Thr Gln Val 690 695 700

Val Pro Gly Val Ala Ala Val Pro Val Glu Glu Glu Thr Thr Ala Val 705 710 715 720

Pro Ser Gly Glu Thr Thr Ala lle Leu Glu Phe Thr Thr Glu Pro Glu
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Asn Gln Thr Glu Trp Glu Pro Ala Tyr Thr Pro Val Gly Thr Ser Pro
740 745 750

Leu Pro Gly Ile Leu Pro Thr Trp Pro Pro Thr Gly Ala Glu Thr Glu

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Glu Pro Ser Pro Ser Glu Val Pro Phe Pro Ser Glu Glu Pro Ser Pro 785 790 795 800

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Phe Pro Ser Glu Glu Pro Phe Pro Ser Lys Glu Pro Ser Pro Ser Glu 1. 820 825 830

Glu Pro Ser Ala Ser Glu Glu Pro Tyr Thr Pro Ser Pro Pro Glu Pro 835 840 845

Ser Trp Thr Glu Leu Pro Ser Ser Gly Glu Glu Ser Gly Ala Pro Asp 850 855 860

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Phe Ser Gly Gln Leu Ser Gly Asp Arg Ala Ser Gly Leu Pro Ser Gly 885 890 895

Asp Leu Asp Ser Ser Gly Leu Thr Ser Thr Val Gly Ser Gly Leu Thr

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905

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Gly Leu Pro Ser Gly Glu Val Leu Glu Thr Thr Ala Pro Gly Val Glu 995 1000 1005

Asp lie Ser Gly Leu Pro Ser Gly Glu Val Leu Glu Thr Thr Ala 1010 1015 1020

Pro Gly Val Glu Asp IIe Ser Gly Leu Pro Ser Gly Glu Val Leu 1025 1030 1035

Glu Thr Thr Ala Pro Gly Val Glu Asp Ile Ser Gly Leu Pro Ser

1040 1045 1050

Gly Glu Val Leu Glu Thr Thr Ala Pro Gly Val Glu Asp Ile Ser 1055 1060 1065

Gly Leu Pro Ser Gly Glu Val Leu Glu Thr Ala Ala Pro Gly Val 1070 1075 1080

Glu Asp I le Ser Gly Leu Pro Ser Gly Glu Val Leu Glu Thr Ala 1085 1090 1095 .

Ala Pro Gly Val Glu Asp IIe Ser Gly Leu Pro Ser Gly Glu Val , ... 1100 1105 1110

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Ala Ala Pro Gly Val Glu Asp lie Ser Gly Leu Pro Ser Gly Glu

1175 1180 1185

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Pro Thr Ala Ser Gly Asp Arg Thr Glu lle Ser Gly Asp Leu Ser 2075 2080 2085

Gly His Thr Ser Gln Leu Gly Val Val IIe Ser Thr Ser IIe Pro 2090 2095 2100

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2120 2125 2130

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Gly Tyr Gly Glu Asp Ala Thr Pro Gly Thr Gly Tyr Thr Gly Leu Ala 115 120 125

Ala lle Gin Leu Pro Lys Lys Ala Gly Asp lle Thr Asn Lys Ala Thr 130 135 140

1. 3

Glu Asn Glu Glu Ser Glu Ala Glu Val Asp Glu Asn Glu Gln Gly Ile 165 170 175

Asn Gly Thr Ser Thr Asn Ser Thr Glu Ala Glu Asn Gly Asn Gly Ser 180 185 190

Ser Gly Gly Asp Asn Gly Glu Glu Glu Glu Glu Glu Ser Val Thr Gly
195 200 . 205

Ala Asn Ala Glu Gly Thr Thr Glu Thr Gly Gly Gln Gly Lys Gly Thr 210 215 220 Ser Lys Thr Thr Ser Pro Asn Gly Gly Phe Glu Pro Thr Thr Pro 225 230 235 240

Pro Gln Val Tyr Arg Thr Thr Ser Pro Pro Phe Gly Lys Thr Thr Thr 245 250 255

Val Glu Tyr Glu Gly Glu Tyr Glu Tyr Thr Gly Val Asn Glu Tyr Asp 260 265 270

Asn Gly Tyr Glu He Tyr Glu Ser Glu Asn Gly Glu Pro Arg Gly Asp 275 280 285

Asn Tyr Arg Ala Tyr Glu Asp Glu Tyr Ser Tyr Phe Lys Gly Gln Gly
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Tyr Asp Gly Tyr Asp Gly Gln Asn Tyr Tyr His His Gln 305 315

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<211> 498

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

⟨222⟩ (19)..(321)

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					•												
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			Ser													•	147
		30		_,_			35		• • •	00,	_,	40	uiu	ury	361	:	
							-					40					
gag	σta	ort o	. 220	242	000	244	040	+	o+ ~	+		.					405
			aag														195
uiu		Vai	Lys	Ar g	Pro		Arg	ıyr	Leu	lyr		irp	Leu	Gly	Ala		•
	45					50					55					٠.	•)
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	Val	Pro	Tyr	Pro	Asp	Pro	Leu	Glu	Pro	Arg	Arg	Glu	Val	Cys	Glu		
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Leu	Asn	Pro	Asp	Cys	Asp	Glu	Leu	Ala	Asp	His	He	Gly	Phe	GIn	Glu		
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gcc	tat	cgg	cgc	ttc	tac	ggc	ccg	gtc	tag	ggtg	tcgc	tc t	gctg	gcct	g		341
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			95					100									
gccg	gcaa	acc d	cagt	tctg	c tc	ctct	ccag	gca	ccct	tct	ttcc	tatt	cc c	ctta	ccctt	-	401
	_		•		,			8					00 0	ortg	00011	•	401
gccc	tgad	ect r	CCSP	ccct	a to	gato	tøøø	gto	CCCS.	tca ·	tooo	aa~+	ac +	0000	aataa		461
J = - U	-5-4	•		, , , , ,	6	o~ - 6	-885	5.00		Lua	-000	ugul	SU L	outid	aa Lää	ı	461
actc	cass	ag s	ppaa	teta	2 22	2222	2200	222	2000								400
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<211> 100

<212> PRT

<213> Homo sapiens

<400> 24

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Giy Ala Ala Phe Val Ser Lys Gin Glu Giy Ser Glu Val Val Lys Arg 🔒 🔞 40 45

Pro Arg Arg Tyr Leu Tyr Gin Trp Leu Giy Ala Pro Vai Pro Tyr Pro 50 55 60

Asp Pro Leu Glu Pro Arg Arg Glu Val Cys Glu Leu Asn Pro Asp Cys 65 70 75 80

Asp Glu Leu Ala Asp His Ile Gly Phe Gln Glu Ala Tyr Arg Arg Phe 85 90 95

Tyr Gly Pro Val 100

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Met Arg Leu Pro Lys Leu Leu Thr Phe Leu Leu	
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Trp Tyr Leu Ala Trp Leu Asp Leu Glu Phe Ile Cys Thr Val Leu Gly	
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Ala Pro Asp Leu Gly Gln Arg Pro Gln Gly Thr Arg Pro Gly Leu Ala	
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aaa gca gag gcc aag gag agg ccc ccc ctg gcc cgg aac gtc ttc agg	496
Lys Ala Glu Ala Lys Glu Arg Pro Pro Leu Ala Arg Asn Val Phe Arg	

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45

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Pro	Gly	Gly	His	Ser	Tyr	Gly	Gly	Gly	Ala	Thr	Asr	n Ala	Asn	Ala	Arg		
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Ala	Lys	Gly	Gly		Gly	Gin	lhr	Gly		Leu	Thr	Gin	Pro		Lys		
				80					85					90			
gat	gaa	CCC	aaa	aag	ctg	CCC	CCC	aga	ccg	ggc	ggc	cct	gaa	ccc	aag		640
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Pro	Gly	His	Pro	Pro	GIn	Thr	Arg	Gln	Ala	Thr	Ala	Arg	Thr	Val	Thr		
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cca	aaa	gga	cag	ctt	ccc	gga	ggc	aag	gca	CCC	сса	aaa	gca	gga	tct		736
Pro		Gly	Gln	Leu	Pro	Gly	Gly	Lys	Ala	Pro	Pro	Lys	Ala	Gly	Ser		
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	Pro	Ser	Ser	Phe		Leu	Lys	Lys	Ala		Glu	Pro	Gly	Pro			
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			,	160			6		165			••••		170	u.u		
tac	atg	ctc	tcg	ctg	tac	agg	acg	ctg	tcc	gat	gct	gac	aga	aag	gga		880
												Asp					
			175					180					185				
ggc	aac	agc	agc	gtg	aag	ttg	gag	gct	ggc	ctg	gcc	aac	acc	atc	acc		928
Gly	Asn	Ser	Ser	Val	Lys	Leu	Glu	Ala	Gly	Leu	Ala	Asn	Thr	lle	Thr		
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Ser			e Asp	Lys	Gly	/ Gin	Asp	Asp	Arg	Gl	y Pro	Va Va	l Val	Ar	g Lys		
	205	5				210	•				215	5					
000		. +															
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220		, iyi	Vai	PHE			ser	Ala	Leu			: Asp	Gly	Let	Leu		
220	,				225)				230)				235		
ggg	gcc	gag	ctg	cgg	ato	ttg	cgg	aag	aag	CCC	tcg:	gac	acg	gcc	aag		1072
															Lys		
				240					245			·		250			
cca	gcg	gcc	ccc	gga	ggc	ggg	cgg	gct	gcc	cag	ctg	aag	ctg	tcc	agc		1120
Pro	Ala	Ala	Pro	Gly	Gly	Gly	Arg	Ala	Ala	Gin	Leu	Lys	Leu	Ser	Ser		
			255					260					265				
																·. ·	•
						ccg									. –		1168
Cys	Pro		Gly	Arg	Gln	Pro	Ala	Ser	Leu	Leu	Asp	Val	Arg	Ser	Va I		
•		270					275					280					
						ggc					•						1216
Pro		Leu	Asp	Gly	Ser	Gly	Trp	Glu	Val	Phe	Asp	He	Trp	Lys	Leu		
	285					290					295						
++0	000	000	+++	000		*											
						tcg								•			1264
300	AI g	ASII	LIIC	LyS		Ser	АІА	uin			Leu	GIU	Leu	Glu			
000					305					310					315		
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gcc	gcc	cgg	cag	gtc	cac	gag	aag	gcc	ctg	ttc	ctg	gtg	ttt	ggc	cgc		1360
						Glu									_		
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Gin	Asp 365	Asp	Lys	Thr	Val	Tyr 370	Glu	Tyr	Leu	Phe	Ser 375	Gin	Arg	Arg	Lys		
												ccc Pro	_				1504
380					385				•	390				•	395	:	
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				400					405					410		1,	š
			Asp					Ala				tac Tyr	Glu	_			1600
	+40	505	415	0+4	+ 00	404	++0	420	++~	000	+00		425	~~~			1640
												cac His 440					1648
acg	aat	cat	gca	gtc	atc	cag	acc	ctg	atg	aac	tcc	atg	gac	ccc	gag		1696
Thr	Asn 445	His	Ala	Val	He	GIn 450	Thr	Leu	Met	Asn	Ser 455	Met	Asp	Pro	Glu		
												agt					1744
Ser 460	ınr	rro	rro	inr	Cys 465	Uys	val	Pro	Inr	Arg 470	Leu	Ser	Pro	He	Ser 475		
			lle	Asp					Val			aag Lys	Gln	Tyr			1792
				480					485					490			

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Asp Met Val Val Glu Ser Cys Gly Cys Arg
495 500

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<210> 26

<211> 501

<212> PRT

<213> Homo sapiens

<400> 26

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Gln Arg Pro Gln Gly Thr Arg Pro Gly Leu Ala Lys Ala Glu Ala Lys 35 40 45

Glu Arg Pro Pro Leu Ala Arg Asn Val Phe Arg Pro Gly Gly His Ser 50 55 60

Tyr Gly Gly Gly Ala Thr Asn Ala Asn Ala Arg Ala Lys Gly Gly Thr 65 70 75 80

Gly Gln Thr Gly Gly Leu Thr Gln Pro Lys Lys Asp Glu Pro Lys Lys

85

90

95

Leu Pro Pro Arg Pro Gly Gly Pro Glu Pro Lys Pro Gly His Pro Pro
100 105 110

Gin Thr Arg Gin Ala Thr Ala Arg Thr Val Thr Pro Lys Gly Gin Leu 115 120 125

Pro Gly Gly Lys Ala Pro Pro Lys Ala Gly Ser Val Pro Ser Ser Phe 130 135 140

Leu Leu Lys Lys Ala Arg Glu Pro Gly Pro Pro Arg Glu Pro Lys Glu 145 150 155 160

Pro Phe Arg Pro Pro Pro IIe Thr Pro His Glu Tyr Met Leu Ser Leu
165 170 175

Tyr Arg Thr Leu Ser Asp Ala Asp Arg Lys Gly Gly Asn Ser Ser Val 180 185 190

Lys Leu Glu Ala Gly Leu Ala Asn Thr Ile Thr Ser Phe Ile Asp Lys
195 200 205

Gly Gin Asp Asp Arg Gly Pro Val Val Arg Lys Gin Arg Tyr Val Phe 210 215 220

Asp lie Ser Ala Leu Glu Lys Asp Gly Leu Leu Gly Ala Glu Leu Arg . 225 230 235 240 . .

Ile Leu Arg Lys Lys Pro Ser Asp Thr Ala Lys Pro Ala Ala Pro Gly
245 250 255

Gly Gly Arg Ala Ala Gln Leu Lys Leu Ser Ser Cys Pro Ser Gly Arg 260 265 270

Gin Pro Ala Ser Leu Leu Asp Val Arg Ser Val Pro Gly Leu Asp Gly 275 280 285

Ser Gly Trp Glu Val Phe Asp lle Trp Lys Leu Phe Arg Asn Phe Lys 290 295 300

Asn Ser Ala Gin Leu Cys Leu Giu Leu Giu Ala Trp Giu Arg Gly Arg 305 310 315 320 Ala Val Asp Leu Arg Gly Leu Gly Phe Asp Arg Ala Ala Arg Gln Val 325 330 335

His Glu Lys Ala Leu Phe Leu Val Phe Gly Arg Thr Lys Lys Arg Asp 340 345 350

Leu Phe Phe Asn Glu IIe Lys Ala Arg Ser Gly Gln Asp Asp Lys Thr . 355 360 365

Val Tyr Glu Tyr Leu Phe Ser Gln Arg Arg Lys Arg Arg Ala Pro Leu 370 375 380

 $r_{r_{i}} \to$

Ala Thr Arg Gin Gly Lys Arg Pro Ser Lys Asn Leu Lys Ala Arg Cys 385 390 395 400

Ser Arg Lys Ala Leu His Val Asn Phe Lys Asp Met Gly Trp Asp Asp 405 410 415

Trp IIe IIe Ala Pro Leu Glu Tyr Glu Ala Phe His Cys Glu Gly Leu
420 425 430

Cys Glu Phe Pro Leu Arg Ser His Leu Glu Pro Thr Asn His Ala Val 435 440 445

lle Gln Thr Leu Met Asn Ser Met Asp Pro Glu Ser Thr Pro Pro Thr 450 455 460 Cys Cys Val Pro Thr Arg Leu Ser Pro Ile Ser Ile Leu Phe Ile Asp 465 470 475 480

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Ser Cys Gly Cys Arg 500

<210> 27

<211> 1378

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> (276).. (1130)

<400> 27

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5

tti	t ggo	tt.	t acg	cag	gag	caa	gtg	gcg	tgo	gtg	tgc	gag	gtt	ct	g cag		341
Phe	e Gly	/ Pho	e Thr	Gln	Glu	Gin	Val	Ala	Cys	Val	Cys	Glu	ı Val	Lei	ı Gin		
			10					15					20				
caa	ggc	gg:	a aac	ctg	gag	cgc	ctg	ggc	agg	tto	ctg	tgg	tca	cte	CCC		389
Gir	Gly	Gly	/ Asn	Leu	Glu	Arg	Leu	Gly	Arg	Phe	Leu	Trp	Ser	Leu	Pro		
		25					30					35					
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gco	tgc	gad	cac	ctg	cac	aag	aac	gag	agc	gta	ctc	aag	gcc	aag	gcg		437
Ala	Cys	Asp	His	Leu	His	Lys	Asn	Glu	Ser	Val	Leu	Lys	Ala	Lys	Ala		
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gtg	gtc	gco	ttc	cac	cgc	ggc	aac	ttc	cgt	gag	ctc	tac	aag	atc	ctg		485
Val	Val	Ala	Phe	His	Arg	Gly	Asn	Phe	Arg	Glu	Leu	Tyr	Lys	He	Leu		
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Glu	Ser	His	Gin	Phe	Ser	Pro	His	Asn	His	Pro	Lys	Leu	Gln	Gln	Leu		
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tgg	ctg	aag	gcg	cat	tac	gtg	gag	gcc	gag	aag	ctg	cgc	ggc	cga	CCC		581
Trp	Leu	Lys	Ala	His	Tyr	Val	Glu	Ala	Glu	Lys	Leu	Arg	Gly	Arg	Pro		
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ctg	ggc	gcc	gtg	ggc	aaa	tat	cgg	gtg	cgc	cga	aaa	ttt	cca	ctg	ccg		629
Leu	Gly	Ala	Val	Gly	Lys	Tyr	Arg	Val	Arg	Arg	Lys	Phe	Pro	Leu	Pro		
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cgc	acc	atc	tgg	gac	ggc	gag	gag	acc	agc	tac	tgc	ttc	aag	gag	aag		677
Arg	Thr	He	Trp	Asp	Gly	Glu	Glu	Thr	Ser	Tyr	Cys	Phe	Lys	Glu	Lys		
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tcg	agg	ggt	gtc	ctg	cgg	gag	tgg	tac .	gcg	cac	aat	ccc	tac	cca	tcg		725
Ser	Arg	Gly	Val	Leu .	Arg	Glu	Trp	Tyr	Ala	His .	Asn	Pro	Tyr	Pro	Ser		
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Pro	Arg	Glu	Lys	Arg	Glu	Leu	Ala	Glu	Ala	Thr	Gly	Leu	Thr	Thr	Thr	
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cag	gtc	agc	aac	tgg	ttt	aag	aac	cgg	agg	caa	aga	gac	cgg	gcc	gcg	821
															Ala	
			170					175				•	180			
																•
gag	gcc	aag	gaa	agg	gag	aac	acc	gaa	aac	aat	aac	tcc	tac	tcc	aac	869
				Arg												000
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		100					130					190				
000	205															047
				ctc										_		917
Lys		ASN	Gin	Leu	Ser		Leu	Glu	Gly	Gly		Pro	Leu	Met	Ser	•
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Ser	Ser	Glu	Glu	Glu	Phe	Ser	Pro	Pro	Gin	Ser	Pro	Asp	GIn	Asn	Ser	
215					220					225					230	
gtc	ctt	ctg	ctg	cag	ggc	aat	atg	ggc	cac	gcc	agg	agc	tca	aac	tat	1013
Val	Leu	Leu	Leu	Gin	Gly	Asn	Met	Gly	His	Ala	Arg	Ser	Ser	Asn	Tyr	
				235					240					245		
tct	ctc	ccg	ggc	tta	aca	gcc	tcg	cag	ccc	agt	cac	ggc	ctg	cag	acc	1061
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			250					255					260			
CAC	Cag	cat	can	ctc	raa	gac	tet	cta	ctc	aac	000	oto	200	tcc	agt	1109
				Leu												1103
1113	um		uiii	Leu	um	voh		LÇU	Leu	uiy	110		1111	961	Set	
		265					270					275				
_4																
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Leu		Asp	Leu	Gly	Ser											
	280															

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<210> 28

<211> 284

<212> PRT

<213> Homo sapiens

<400> 28

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Phe Leu Trp Ser Leu Pro Ala Cys Asp His Leu His Lys Asn Glu Ser 35 40 45

Val Leu Lys Ala Lys Ala Val Val Ala Phe His Arg Gly Asn Phe Arg 50 55 60

Glu Leu Tyr Lys IIe Leu Glu Ser His Gln Phe Ser Pro His Asn His 65 70 75 80 Pro Lys Leu Gln Gln Leu Trp Leu Lys Ala His Tyr Val Glu Ala Glu 85 90 95

Lys Leu Arg Gly Arg Pro Leu Gly Ala Val Gly Lys Tyr Arg Val Arg
100 105 110

Arg Lys Phe Pro Leu Pro Arg Thr lle Trp Asp Gly Glu Glu Thr Ser 115 120 125

Tyr Cys Phe Lys Glu Lys Ser Arg Gly Val Leu Arg Glu Trp Tyr Ala 130 135 140

His Asn Pro Tyr Pro Ser Pro Arg Glu Lys Arg Glu Leu Ala Glu Ala 145 150 155 160

Thr Gly Leu Thr Thr Gln Val Ser Asn Trp Phe Lys Asn Arg Arg
165 170 175

Gin Arg Asp Arg Ala Ala Giu Ala Lys Giu Arg Giu Asn Thr Giu Asn 180 185 190

Asn Asn Ser Ser Asn Lys Gln Asn Gln Leu Ser Pro Leu Glu Gly
195 200 205

Gly Lys Pro Leu Met Ser Ser Glu Glu Glu Phe Ser Pro Pro Gln 210 215 220 Ser Pro Asp Gln Asn Ser Val Leu Leu Leu Gln Gly Asn Met Gly His 225 230 235 240

Ala Arg Ser Ser Asn Tyr Ser Leu Pro Gly Leu Thr Ala Ser Gln Pro
245 250 255

Ser His Gly Leu Gln Thr His Gln His Gln Leu Gln Asp Ser Leu Leu 260 265 • 270

Gly Pro Leu Thr Ser Ser Leu Val Asp Leu Gly Ser 275 280

<210> 29

<211> 2590

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> (546).. (1112)

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<221> CDS

<222> (2036).. (2071)

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Leu Gin Gly Ala Arg Arg Ala Gly Gly Arg Arg Ala Gly Gly Gly 50 55 60

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Ala Leu Arg Thr Leu IIe Pro Thr Glu Pro Ala Asp Arg Lys Leu Ser 100 105 110

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Gin lie Cys Thr Phe Cys Leu Ser Asn Gin Arg Lys Leu Ser Lys Asp 180 185 190

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